17/02/2005 10081147

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             JANUARY 10 CURRENT WINDOWS VERSION IS V7.01a, CURRENT
              MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
              AND CURRENT DISCOVER FILE IS DATED 10 JANUARY 2005
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Page 1 SAEED

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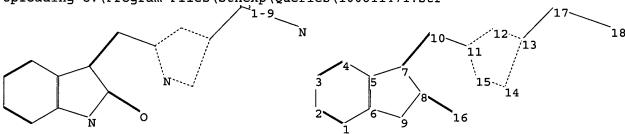
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chain nodes : 10 16 17 1

ring nodes :

1 2 3 4 5 6 7 8 9 11 12 13 14 15

17/02/2005 10081147

chain bonds :

7-10 8-16 10-11 13-17 17-18

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-9 7-8 8-9 11-12 11-15 12-13 13-14 14-15

exact/norm bonds :

5-7 6-9 7-8 8-9 8-16 11-12 11-15 12-13 13-14 14-15 17-18

exact bonds : 7-10 10-11 13-17 normalized bonds :

1-2 1-6 2-3 3-4 4-5 5-6

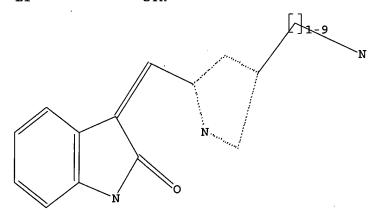
Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:CLASS 17:CLASS 18:CLASS

## L1 STRUCTURE UPLOADED

=> d L1 HAS NO A

L1 HAS NO ANSWERS L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> s 11

SAMPLE SEARCH INITIATED 11:12:15 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 175 TO ITERATE

100.0% PROCESSED 175 ITERATIONS

25 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*
BATCH \*\*COMPLETE\*\*

PROJECTED ITERATIONS: 2707 TO 4293 PROJECTED ANSWERS: 200 TO 800

L2 25 SEA SSS SAM L1

=> s l1 full

Page 3

SAEED

17/02/2005 10081147

FULL SEARCH INITIATED 11:12:32 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 4152 TO ITERATE

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591 ANSWERS

SEARCH TIME: 00.00.01

L3 591 SEA SSS FUL L1

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=> s 13

L4 65 L3

=> d ibib abs hitstr tot

## 10081147

L4 ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2005:2190 CAPLUS
DOCUMENT NUMBER: 142:93676
A preparation of sulfonamide substituted indolinones, useful as inhibitors of DNA dependent protein kinase (DNA-PK)
INVENTOR(S): Howlett, Anthony R.; Rice, Audie; Moshinsky, Deborsh; Hammarsten, Ola Sugen, Inc., USA U.S. Pat. Appl. Publ., 46 pp. CODEN: USX/COD DOCUMENT TYPE: Patent English
FAMILY ACC. NUM. COUNT: 1

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| PATENT NO.             | KIND | DATE     | APPLICATION NO.   | DATE     |
|------------------------|------|----------|-------------------|----------|
|                        |      |          |                   |          |
| US 2004266843          | A1   | 20041230 | US 2004-793943    | 20040308 |
| PRIORITY APPLN. INFO.: |      |          | US 2003-452549P P | 20030307 |

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB The invention relates to a preparation of sultonamius subsection indolinones of formula I (wherein: R1 and R2 are independently selected from H, (un)substituted Ph, thiazolyl, or alkyl, etc.: R3, R4, and R5 are independently selected from H or alkyl], useful as inhibitors of DNA dependent protein kinase (DNA-PK). The invention relates to the field of radiosensitizing agents which are capable of enhancing radiotherapy by inhibiting DNA-PK (DNA-Protein kinase). For instance, sulfonamide aubstituted indolinone II was prepared via condensation of pyrrole derivative

derivative
III and indole derivative IV. The prepared indolinone derivative V was found to

d to inhibit DNA-PK (IC50 = 1.6 μM). 342641-63-89 775321-58-99 775321-59-09 775321-60-39 775321-68-19 775321-69-29 775321-75-09 775321-75-09 775321-77-29 775321-77-09 775321-77-29 775321-78-39 775321-73-09 775321-77-29 775321-78-39 775322-07-19 775322-08-29 775322-13-99 775322-13-39 77532-

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(USES) (preparation of sulfonamide substituted indolinones useful as inhibitors of DNA dependent protein kinase (DNA-PK))

ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) PAGE 1-A

PAGE 1-B

775321-60-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-[5-[(3,4-dihydro-1{2H})-quinolinyl) sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[3-(4-methyl-1-piperazinyl)propyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

775321-68-1 CAPLUS
1H-Pyrrole-3-acetamide, 5-[(2)-[5-[(2,3-dihydro-1H-indol-1-yl)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 342641-63-8 CAPLUS H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-(2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

775321-58-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(Z)-{5-{(3,4-dihydro-2(1H)-isoquinolinyl)sulfonyl}-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-{3-(4-methyl-1-piperazinyl)propyl}- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

PAGE 1-A

PAGE 1-B

Double bond geometry as shown.

ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

775321-69-2 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(z)-[5-[(z)-3-dihydro-5-methoxy-1H-indol-1-yl)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9C1) (CA INDEX NAME)

Double bond geometry as shown.

PAGE 1-B

-NEt2

RN 775321-70-5 CAPLUS
CN 1H-Pyrrole-3-acetamide,
N-[2-[diethylamino]ethyl]-5-[[Z]-[5-[[Z],3-dihydro1H-indol-1-yl]]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

Double bond geometry as shown.

775321-75-0 CAPLUS
1H-Pyrrole-3-acetamide, 5-[{Z}-[5-[{2,3-dihydro-1H-indol-1-y1}sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown. ,

775321-77-2 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-[5-[(Z,3-

ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 775321-81-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5<sup>2</sup>[(2)-{5-((2,3-dihydro-1H-indol-1-yl)sulfonyl}-

1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl)-2,4-dimethyl-N-[3-(4-methyl-1-piperazinyl)propyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

PAGE 1-A

PAGE 1-B

775322-06-0 CAPLUS 1H-Pyrrole-3-acetamide, 5-{(2)-[5-(aminosulfonyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Contin dihydro-1H-indol-1-yl)sulfonyl|-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl|-2,4-dimethyl- (9CI) (CA INDEX NAME) (Continued)

Double bond geometry as shown.

RN 775321-78-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-((2)-(5-[(2,3-dihydro-1H-indol-1-yl)sulfonyl)1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl)-2,4-dimethyl-N-(2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 775321-80-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[(2,3-dih)ydro-1H-indol-1-yl)sulfonyl]1,2-dihydro-2-xoo-3H-indol-3-ylidene|methyl]-2-ethyl-4-methyl-N-[2-[1-yyrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

775322-07-1 CAPLUS
1H-Pyrrole-3-acetamide, 5-{(Z)-[1,2-dihydro-5-{(methylamino)sulfonyl]-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(l-pyrrolidinyl)ethyl]-(9C) (CA INDEX NAME)

Double bond geometry as shown.

RN 775322-08-2 CAPLUS
CN 1H-Pyrrole-3-acetamide,
5-[(2)-[5-[(dimethyl]amino) sulfonyl]-1,2-dihydro-2oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl](9C1) (CA INDEX NAME)

Double bond geometry as shown.

RN 775322-12-8 CAPLUS
CN 1H-Pyrrole-3-acetamide,
5-{(2)-{5-[(3-chlorophenyl)methylamino]sulfonyl}1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl}-2,4-dimethyl-N-[2-{1pyrrolidinyl}ethyl]- (9CI) (CA INDEX NAME)

Page 6

SAEED

L4 ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) Double bond geometry as shown.

 $\label{eq:capture} \raiseta 2.2-13-9 \quad CAPLUS \\ $1$H-Pyrrole-3-acetamide, $5-[(Z)-[5-(aminosulfonyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) \\ \raiseta 2.2-dimethyl-1-2-(diethylamino)ethyl]-2,4-dimethyl-1-2-(diethylamino)ethyl-1-2-(diethylamin$ 

(CA INDEX NAME)

Double bond geometry as shown.

775322-14-0 CAPLUS IH-Pyrrole-3-acetamide, N-{2-(diethylamino)ethyl}-5-{(Z)-[1,2-dihydro-5-(methylamino)sulfonyl}-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl-(SCI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

775322-19-5 CAPLUS 1H-Pyrrole-3-acetamide, 5-[(2)-[1,2-dihydro-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 775322-20-8 CAPLUS
CN 1H-Pyrrole-3-acetamide,
5-[(2]-[5-[(dimethylamino)sulfonyl]-1,2-dihydro-2oxo-3M-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl](9CI) (CA INDEX NAME)

Double bond geometry as shown.

775322-25-3 CAPLUS
1H-Pyrrole-3-acetamide, N-[2-(diethylamino)ethyl]-5-[{Z}-[5-

[(dimethylamino)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

775322-15-1 CAPLUS
1H-Pyrrole-3-acetamide, 5-[(Z)-[5-(aminosulfonyl)-1,2-dihydro-2-oxo-3H-indol-3-yildene]methyl}-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

RN 775322-18-4 CAPLUS
CN 1H-Pyrrole-3-acetamide,
5-[(Z)-(5-{((3-chlorophenyl)methylamino)sulfonyl}-

1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl[-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

819044-92-3 CAPLUS
1H-Pyrrole-3-carboxamide, N-(3-ethylpentyl)-5-[(Z)-[5-[(5-fluoro-2,3-dihydro-1+-indol-1-yl)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

PAGE 1-B

PAGE 1-A

-CHEt2

L4 ANSWER 2 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2004:1121013 CAPLUS

TITLE: Synergistic effect of SU11248 with cytarabine or daunorubicin on FLT3 ITD-positive leukemic cells

Yee, Kevin W. H.; Schittenhelm, Marcus; O'Farrell, Anne-Marie; Town, Ajia R.; McGreevey, Laura; Bainbridge, Troy; Cherrington, Julie M.; Heinrich, McCheel C.

CORPORATE SOURCE: Oregon Health and Science University Cancer Institute and Portland, OR, USA

SOURCE: Blood (2004), 104(13), 4202-4209

CODEN: BLOODW; ISSN: 0006-4971

PUBLISHER: American Society of Hematology

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Fetal liver tyrosine Kinase 3 internal tandem duplication (FLT3 ITD)

mutations are the most common mol. abnormality associated with adult acute

myeloid leukemia (AML). To exploit this mol. target, a number of potent

and

specific FLT3 kinase inhibitors have been developed and are currently
being tested in early phase clin. triels of patients with refractory AML.

To explore the efficacy of combining a FLT3 inhibitor with standard AML
chemotherapy drugs, we tested the effect of combining the FLT3 inhibitor
SU11248 with cyterabine or daunorubicin on the proliferation and survival
of cell lines expressing either mutant (FLT3 TTD or FLT3 D835V) or
wild-type (WT) FLT3. SU11248 had additive-to-synergistic inhibitory
effects on FLT3-dependent leukemic cell proliferation when combined with
cytarabine or daunorubicin. The synergistic interaction of SU11248 and
the traditional antileukemic spents was more pronounced for induction of
apoptosis. SU1248 inhibited the proliferation of primary AML
myeloblasts
expressing mutant FLT3 ITD but not WT FLT3 protein. Combining SU11248

expressing mutant FLT3 ITD but not WT FLT3 protein. Combining SU11248

cytarabine synergistically inhibited the proliferation of primary AML myeloblasts expressing FLT3 ITD but not WT FLT3 protein. These data suggest that the addition of potent FLT3 inhibitors such as SU11248 to

AML

chemotherapy regimens could result in improved treatment results.

557795-19-4, SU 11248

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological atudy); USES (Usea) (aymergistic effect of SU11248 with cytarabine or daunorubicin on FLT3 ITD-pos. leukemic cells)

557795-19-4 CaPLUS

1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1, 2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 3 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2004:1059176 CAPLUS
DOCUMENT NUMBER: 142:32986

INVENTOR(S): 101bibitor for the treatment of diabetes
Hagerkvist, Robert Per: Welsh, Nils Richard
Swed.
DOCUMENT TYPE: PT Int. Appl., 22 pp.
CODEN: PIXXD2
DOCUMENT TYPE: PARLLY ACC. NUM. COUNT: 1

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| PAT      | ENT  | NO.  |      |     | KIN | D   | DATE |      |     | APPL | ICAT | ION I | NO. |     | D.  | ATE  |     |
|----------|------|------|------|-----|-----|-----|------|------|-----|------|------|-------|-----|-----|-----|------|-----|
|          |      |      |      |     |     | -   |      |      |     |      |      |       |     |     | _   |      |     |
| WO       | 2004 | 1057 | 63   |     | A2  |     | 2004 | 1209 | 1   | WO 2 | 004- | EP56  | 79  |     | 2   | 0040 | 526 |
|          | W:   | ΑE,  | AG,  | AL, | AM, | ΑT, | ΑU,  | AZ,  | BA, | BB,  | BG,  | BR,   | BW, | BY, | BZ, | CA,  | CH, |
|          |      | CN,  | CO,  | CR, | CU, | CZ, | DE,  | DK,  | DM, | DZ,  | EC,  | ΕE,   | EG, | ES, | FI, | GB,  | GD, |
|          |      | GΕ,  | GH,  | GM, | HR, | ΗU, | ID,  | IL,  | IN, | ÍS,  | JP,  | KE,   | KG, | ΚP, | KR, | ΚZ,  | LC, |
|          |      | LK,  | LR,  | LS, | LT, | LU, | LV,  | ΜA,  | MD, | MG,  | ΜK,  | MN,   | MW, | MX, | MZ, | NA,  | NI, |
|          |      | NO,  | ΝZ,  | OM, | ₽G, | PH, | PL,  | PT,  | RO, | RU,  | SC,  | SD,   | SE, | SG, | SK, | SL,  | SY, |
|          |      | ТJ,  | TM,  | TN, | TR, | TT, | ΤZ,  | UA,  | UG, | us,  | UZ,  | VC,   | VN, | YU, | ZA, | ZM,  | ZW  |
|          | RW:  | BW,  | GH,  | GM, | KE, | LS, | MW,  | MZ,  | NA, | SD,  | SL,  | SZ,   | TZ, | UG, | ZM, | ZW,  | AM, |
|          |      | ΑZ,  | BY,  | KG, | ΚZ, | MD, | RU,  | TJ,  | TM, | ΑT,  | BE,  | BG,   | CH, | CY, | CZ, | DE,  | DK, |
|          |      | EE,  | ES,  | FI, | FR, | GB, | GR,  | Hυ,  | ΙE, | IT,  | LU,  | MC,   | NL, | PL, | PT, | RO,  | SE, |
|          |      | SI,  | sĸ,  | TR, | BF, | ₿J, | CF,  | CG,  | CI, | CM,  | GΑ,  | GN,   | GQ, | GW, | ML, | MR,  | ΝE, |
|          |      | SN,  | TD,  | TG  |     |     |      |      |     |      |      |       |     |     |     |      |     |
| PRIORITY | APP  | LN.  | INFO | .:  |     |     |      |      | -   | GB 2 | 003- | 1208  | 6   |     | A 2 | 0030 | 527 |

GB 2003-12086 A 20030527

AB The invention discloses the use of a c-Abl-, PDGFR-, or c-kit-tyrosine kinase inhibitor, e.g. I, or a pharmaceutically acceptable salt thereof, for the manufacture of a medicament for the treatment of diabetes, including

type I or type II diabetes.

1 53795-19-4, 5U 11248

RI: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(SU 11248; c-abl-, PDGFR-, or c-kit-tyrosine kinase inhibitor for

ANSWER 2 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

REFERENCE COUNT:

51 THERE ARE 51 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

10081147

L4 ANSWER 4 OF 65 CAPLUS COPYRIGHT 2005 ACS ON STN ACCESSION NUMBER: 2004:995713 CAPLUS DOCUMENT NUMBER: 141:420610 TITLE: Surface reconstructions Surface receptor complexes as biomarkers of disease and for determination of treatment with dimer-acting

drugs Chan-Hui, Po-Ying; Dua, Rajiv; Mukherjee, Ali; Pidaparthi, Sailaja; Salimi-Moosavi, Hossein; Shi, Yining; Singh, Sharat INVENTOR (S):

PATENT ASSIGNEE(S): SOURCE: USA
U.S. Pat. Appl. Publ., 67 pp., Cont.-in-part of U.S. Ser. No. 623,057.
CODEN: USXXCO

DOCUMENT TYPE: LANGUAGE: Patent

FAMILY ACC. NUM. COUNT:

| PATENT INFORMATION.    |      |          |                 |    |          |
|------------------------|------|----------|-----------------|----|----------|
| PATENT NO.             | KIND | DATE     |                 |    | DATE     |
| US 2004229293          |      | 20041118 | US 2004-812619  |    | 20040330 |
| US 2003013126          | A1   | 20030116 | US 2002-154042  |    | 20020521 |
| US 2004126818          | A1   | 20040701 | US 2003-623057  |    | 20030717 |
| US 2004197835          | A1   | 20041007 | US 2004-830543  |    | 20040422 |
| PRIORITY APPLN. INFO.: |      |          | US 2002-154042  | A2 | 20020521 |
|                        |      |          | US 2002-398724P | P  | 20020725 |
|                        |      |          | US 2003-459888P | P  | 20030401 |
|                        |      |          | US 2003-623057  | A2 | 20030717 |
|                        |      |          | US 2003-494482P | P  | 20030811 |
|                        |      |          | US 2003-508034P | P  | 20031001 |
|                        |      |          | US 2003-512941P | P  | 20031020 |
|                        |      |          | US 2003-523258P | P  | 20031118 |
|                        |      |          | US 2001-292548P | P  | 20010521 |
|                        |      |          | US 2001-334901P | P  | 20011024 |

AB The invention is directed to a new class of biomarker in patient samples comprising dimers of cell surface membrane receptors. In one aspect, th invention includes a method of determining the status of a disease or healthful

condition by correlating such condition to amts. of one or more dimers of cell surface membrane receptors measured directly in a patient sample, in particular a fixed tissue sample. In another aspect, the invention includes a method of determining a status of a cancer in a specimen from

individual by correlating measurements of amts. of one or more dimers of cell surface membrane receptors in cells of the specimen to such status, including presence or absence of a pre-cancerous state, presence or

L4 ANSWER 5 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN ACCESSION NUMBER: 2004:965067 CAPLUS DOCUMENT NUMBER: 141:406039

141:406039
Combinations for the treatment of diseases involving cell proliferation, migration or apoptosis of myeloma cells, or angiogenesis Hilberg, Frank; Solca, Flavio; Stefanic, Martin Friedrich; Baum, Anke; Munzert, Gerd; Van Meel, Jacobus C. A.
Boehringer Ingelheim International G.m.b.H., Germany; Boehringer Ingelheim Pharma G.m.b.H. & Co. K.-G. PCT Int. Appl., 101 pp.
CODEN: PIXXD2
Patent TITLE:

INVENTOR (S):

PATENT ASSIGNEE (S):

SOURCE:

Patent

DOCUMENT TYPE: LANGUAGE: English 2

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PR

|       |       |       |      |     |     | _   |      |      |     |      |      |      |     |     | _   |      |     |
|-------|-------|-------|------|-----|-----|-----|------|------|-----|------|------|------|-----|-----|-----|------|-----|
|       |       | NO.   |      |     |     |     | DATE |      |     |      |      |      |     |     |     |      |     |
|       |       |       |      |     |     |     |      |      |     |      |      |      |     |     |     |      |     |
| WO    | 2004  | 10962 | 24   |     | A2  |     | 2004 | 1111 | 1   | WO 2 | 004- | EP43 | 63  |     | 2   | 0040 | 424 |
| WO    | 2004  | 10962 | 24   |     | A3  |     | 2004 | 1216 |     |      |      |      |     |     |     |      |     |
|       | W:    | ΑE,   | AG,  | AL, | AM, | ΑT, | AU,  | ΑZ,  | BA, | BB,  | BG,  | BR,  | BW, | BY, | BZ, | CA,  | CH, |
|       |       | CN,   | co,  | CR, | CU, | CZ, | DK,  | DM,  | DZ, | EC,  | EE,  | EG,  | ES, | FI, | GB, | GD,  | GE, |
|       |       | GH,   | GM,  | HR, | ΗU, | ID, | IL,  | IN,  | IS, | JP,  | KE,  | KG,  | KP, | KR, | KZ, | LC,  | LK, |
|       |       | LR,   | LS,  | LT, | LU, | LV, | MA,  | MD,  | MG, | MK,  | MN,  | MW,  | MX, | MZ, | NA, | NI,  | NO, |
|       |       | NZ,   | OM,  | PG, | PH, | PL, | PT,  | RO,  | RU, | SC,  | SD,  | SE,  | SG, | SK, | SL, | SY,  | TJ, |
|       |       | TM,   | TN,  | TR, | TT, | TZ, | UA,  | UG,  | US, | UZ,  | VC,  | VN,  | YU, | ZA, | ZM, | ZW   |     |
|       | RW:   | BW,   | GH,  | GM, | KΕ, | LS, | MW,  | MZ,  | NA, | SD,  | SL,  | SZ,  | TZ, | UG, | ZM, | ZW,  | AM, |
|       |       | AZ,   | BY,  | KG, | KZ, | MD, | RU,  | TJ,  | TM, | AT,  | BE,  | BG,  | CH, | CY, | CZ, | DE,  | DK, |
|       |       | EE,   | ES,  | FI, | FR. | GB, | GR,  | HU,  | IE, | IT,  | LU,  | MC,  | NL, | PL, | PT, | RO,  | SE, |
|       |       | SI,   | SK,  | TR, | BF, | BJ, | CF,  | CG,  | CI, | CM,  | GΑ,  | GN,  | GQ, | G₩, | ML, | MR,  | NE, |
|       |       | SN,   | TD,  | TG  |     |     |      |      |     |      |      |      |     |     |     |      |     |
| EP    | 1473  | 3043  |      |     | A1  |     | 2004 | 1103 |     | EP 2 | 003- | 9587 |     |     | 21  | 0030 | 429 |
|       | R:    | AT,   | BE,  | CH, | DE, | DK, | ES,  | FR,  | GB, | GR,  | IT,  | LI,  | LU, | NL, | SE, | MC,  | PT, |
|       |       | IE,   | SI,  | LT, | LV, | FI, | RO,  | MK,  | CY, | AL,  | TR,  | BG,  | CZ, | EE, | HU, | SK   |     |
| IORIT | Y APE | LN.   | INFO | . : |     |     |      |      | 1   | EP 2 | 003- | 9587 |     | 4   | A 2 | 0030 | 429 |
|       |       |       |      |     |     |     |      |      | 1   | EP 2 | 004- | 508  |     | 1   | A 2 | 0040 | 113 |
|       |       |       |      |     |     |     |      |      | 1   | EP 2 | 004- | 1171 |     | 1   | A 2 | 0040 | 121 |

The present invention relates to a pharmaceutical combination for the treatment of diseases which involves cell proliferation, migration or apoptosis of myeloma cells, or angiogenesis. The invention also relates to a method for the treatment of said diseases, comprising co-administration of effective amts. of specific active compds. and/or co-treatment with radiation therapy, in a ratio which provides an

and synergistic effect, and to the combined use of these specific compds. and/or radiotherapy for the manufacture of corresponding pharmaceutical combination preprise. The pharmaceutical combination can include selected protein tyrosine kinase receptor antagonists and further chemotherapeutic or naturally occurring semisynthetic or synthetic agents. 357785-19-4 additive

537793-19-4
RI: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
(Biological study); USES (Uses)
(SU 11248; drug combinations for diseases involving cell proliferation and migration or apoptosis or anglogenesis including protein tyrosine kinase receptor antagonists and radiotherapy)

ANSWER 4 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) absence of a cancerous state, prognosis of a cancer, or responsiveness to treatment. Preferably, methods of the invention are implemented by using sets of binding components of one or more types of receptor dimers. After binding, mol. tags are released and sepd. from the assay mixt. for anal. 557795-19-4, SU11248
RL: PAC (Pharmacological activity): THU (Therapeutic use): BIOL (Biological atudy): USES (Uses) (surface receptor complexes as biomarkers of disease or responsiveness to treatment)
SH-Pytrole-3-carboxamide, N-(2-(diethylamino)ethyl]-5-[{2}-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 5 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
557795-19-4 CAPLUS
H-Pyrrole-3-carboxamide, N-{2-(diethylamino)ethyl}-5-[{2}-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl- (9CI) (CA INDEX

Double bond geometry as shown.

627908-92-3
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Blological study); USES (Uses)
(SU 14613; drug combinations for diseases involving cell proliferation and migration or apoptosis or angiogenesis including protein tyrosine kinase receptor antagonists and radiotherapy)
627908-92-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[(ZS)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

Page 9 SAEED

#### 10081147

L4 ANSWER 6 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
DOCUMENT NUMBER:
111:361107
Methods for the detection of cell surface receptor complexes as cancer blomarkers and therapeutic effectiveness of cleavage thereof
Chan-Hul, Po-Ying; Salimi-Moosavi, Hossein; Shi, Yining; Singh, Sharat; Dua, Rajiv; Mukherjee, Ali; Pidaparthi, Saliaja
Aclare Biosciences, Inc., USA
POTINENT TYPE:
DOCUMENT TYPE:
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
20
PATENT INFORMATION:
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21
204:905883 CAPLUS
Cleavage Aclare
Chan-Hul, Po-Ying; Salimi-Moosavi, Hossein; Shi, Yining; Singh, Sharat; Dua, Rajiv; Mukherjee, Ali;
Pidaparthi, Saliaja
Aclare Biosciences, Inc., USA
PCT Int. Appl., 106 pp.
COODE: PIXXD2
Patent InfoRMATION:
22
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204:90583 CAPLUS
204:90583 CAPLUS
204:905883 CAPLUS
205:905883 CAPLUS
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205:905883 CAPL

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

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|       |       |      |      |     |     |     |      |      |     |       | ICAT  |       |     |     |     |      |     |
|-------|-------|------|------|-----|-----|-----|------|------|-----|-------|-------|-------|-----|-----|-----|------|-----|
|       |       |      |      |     |     |     |      |      |     |       |       |       |     |     |     |      |     |
| WO    | 2004  | 0923 | 53   |     | A2  |     | 2004 | 1028 | ,   | WO 2  | 2004- | US 97 | 17  |     | 2   | 0040 | 330 |
|       | W:    | ΑE,  | AG,  | AL, | AM, | AT, | AU,  | AZ,  | BA, | BB,   | BG,   | BR,   | BW, | BY, | ΒZ, | CA,  | CH, |
|       |       | CN,  | co,  | CR, | CU, | CZ, | DE,  | DK,  | DM, | DZ,   | EC,   | EE,   | EG, | ES, | FI, | GB,  | GD, |
|       |       | GE,  | GH,  | GM, | HR, | HU, | ID,  | IL,  | IN, | IS,   | JP,   | ΚE,   | KG, | KP, | KR, | ΚZ,  | LC, |
|       |       | LK,  | LR,  | LS, | LT, | LU, | LV,  | MA,  | MD, | MG,   | MK,   | MN,   | MW, | MX, | MZ, | NΑ,  | NI, |
|       |       | NO.  | NZ.  | OM, | PG, | PH, | PL,  | PT,  | RO, | RU,   | SC,   | SD,   | SE, | SG, | SK, | SL,  | SY, |
|       |       | TJ.  | TM.  | TN. | TR. | TT. | TZ,  | UA,  | UG, | US.   | UZ,   | VC,   | VN, | YU, | ZA, | ZM,  | ZW  |
|       | RW:   | BW.  | GH.  | GM. | KE. | LS. | MW.  | MZ.  | SD, | SL.   | SZ,   | TZ.   | UG, | ZM, | ZW, | AM,  | AZ, |
|       |       | BY.  | KG.  | KZ. | MD. | RU, | TJ.  | TM.  | AT. | BE.   | BG,   | CH.   | CY, | CZ. | DE, | DK,  | EE, |
|       |       | ES.  | FI.  | FR, | GB, | GR, | HU,  | IE,  | IT, | LU,   | MC,   | NL,   | PL, | PT, | RO, | SE,  | SI, |
|       |       |      |      |     |     |     |      |      |     |       | GN,   |       |     |     |     |      |     |
|       |       | TD.  |      |     |     |     | •    |      |     |       |       |       |     |     |     |      |     |
| us    | 2004  | 1268 | 18   |     | A1  |     | 2004 | 0701 |     | US 2  | 003-  | 6230  | 57  |     | 2   | 0030 | 717 |
| CORIT | Y APP | LN.  | INFO | . : |     |     |      |      |     | US 2  | 003-  | 4598  | 88P |     | P 2 | 0030 | 401 |
|       |       |      |      |     |     |     |      |      |     |       |       |       |     |     |     |      |     |
|       |       |      |      |     |     |     |      |      | 1   | US 2  | 003-  | 6230  | 57  | i   | A 2 | 0030 | 717 |
|       |       |      |      |     |     |     |      |      | 1   | US 2  | 003-  | 4944  | 82P |     | P 2 | 0030 | 811 |
|       |       |      |      |     |     |     |      |      |     |       |       |       |     |     |     |      |     |
|       |       |      |      |     |     |     |      |      |     | US 2  | 003-  | 5080  | 34P |     | P 2 | 0031 | 001 |
|       |       |      |      |     |     |     |      |      | 1   | US 2  | 003-  | 5129  | 41P |     | P 2 | 0031 | 020 |
|       |       |      |      |     |     |     |      |      |     | 11e 2 | 003-  | 5222  | 50D |     |     | 0031 | 110 |
|       |       |      |      |     |     |     |      |      |     | 03 2  | .003- | JLJE  | JUE |     |     | 0031 | 110 |

The invention is directed to a new class of biomarker in patient samples comprising dimers of cell surface membrane receptors. In one aspect, the invention includes a method of determining the status of a disease or

US 2002-398724P

invention includes a method of determining the state of one or more dimers of condition by correlating such condition to amts. of one or more dimers of cell surface membrane receptors measured directly in a patient sample, in particular a fixed tissue sample. In another aspect, the invention includes a method of determining a status of a cancer in a specimen from

individual by correlating measurements of amts. of one or more dimers of

L4 ANSWER 7 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2004:902199 CAPLUS
DOCUMENT NUMBER: 141:374704

Composition and uses of galectin antagonists to augment treatment of cancer or other proliferative disorders

INVENTOR(S): Chang, Yan; Sasak, Vodek
Glycogenesys, Inc., USA
FOR INC., NUM. COUNT: PIXXD2

DOCUMENT TYPE: PARTIX ACC. NUM. COUNT: 3

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| 1    | PAT | ENT  | NO.                             |   |   | KIN   |   | DATE  |   |   |   |                                 |   |   |   | D   | ATE   |  |  |
|------|-----|------|---------------------------------|---|---|---|---|---|---|---|---|---------------------------------|---|---|---|---|---|--|--|
| ,    | 70  | 200  | 10916                           | 34  |   |   |   | 2004  | 1028  |   |   | 004-                            |   |   |   | 2   | 0040  | 407  |  |
|      |     |      | GE,<br>LK,<br>NO,<br>TJ,<br>BW, | CO,<br>GH,<br>LR,<br>NZ,<br>TM,<br>GH,<br>KG, | CR,<br>GM,<br>LS,<br>OM,<br>TN,<br>GM,<br>KZ, | CU,<br>HR,<br>LT,<br>PG,<br>TR,<br>KE,<br>MD, | CZ,<br>HU,<br>LU,<br>PH,<br>TT,<br>LS,<br>RU, | DE,<br>ID,<br>LV,<br>PL,<br>TZ,<br>MW,<br>TJ, | DK,<br>IL,<br>MA,<br>PT,<br>UA,<br>MZ,<br>TM, | DM,<br>IN,<br>MD,<br>RO,<br>UG,<br>SD,<br>AT, | DZ,<br>IS,<br>MG,<br>RU,<br>US,<br>SL,<br>BE, | EC,<br>JP,<br>MK,<br>SC,<br>UZ, | EE,<br>KE,<br>MN,<br>SD,<br>VC,<br>TZ,<br>CH, | EG,<br>KG,<br>MW,<br>SE,<br>VN,<br>UG,<br>CY, | ES,<br>KP,<br>MX,<br>SG,<br>YU,<br>ZM,<br>CZ, | FI,<br>KR,<br>MZ,<br>SK,<br>ZA,<br>ZW,<br>DE, | GB,<br>KZ,<br>NA,<br>SL,<br>ZM,<br>AM,<br>DK, | GD,<br>LC,<br>NI,<br>SY,<br>ZW<br>AZ,<br>EE, |  |
|      |     |      |                                 | TR,   |   |   |   |   |   |   |   | GN,                             |   |   |   |   |   |  |  |
|      | JS  | 2004 | 10239                           | 25  |   | A1  |   | 2004  | 0205  | 1   | US 2  | 003-                            | 4087  | 23  |   | 2   | 0030  | 407  |  |
| RIOR |     |      | 12239<br>PLN.                   |   |   |   |   |   | 1111  |   |   | 004-<br>003-                    |   |   |   |   |   |  |  |
|      |     |      |                                 |   |   |   |   |   |   | 1   | US 2  | 003-                            | 4610  | 06P   | 1   | P 2   | 0030  | 407  |  |
|      |     |      |                                 |   |   |   |   |   |   | 1   | US 2  | 003~                            | 4745  | 62 P  | 1   | P 2   | 0030  | 530  |  |
|      |     |      |                                 |   |   |   |   |   |   | 1   | US 2  | 001-                            | 2999  | 91P   | 1   | P 2   | 0010  | 621  |  |

The present invention is directed to methods and compns. for augmenting treatment of cancers and other proliferative disorders. In particular embodiments, the invention combines the administration of an agent that inhibits the anti-apoptotic activity of galectin-3 (e.g., a 'galectin-3 inhibitor') so as to potentiate the toxicity of a chemotherapeutic agent. In certain preferred embodiments, the conjoint therapies of the present invention can be used to improve the efficacy of those chemotherapeutic agents whose cytotoxicity is influenced by the status of an -apoptotic Bcl-2 protein for the treated cell. For instance, galectin-3 inhibitors can be administered in combination with a chemotherapeutic agent that interferes with DNA replication fidelity or cell-cycle progression of cells undergoing unwanted proliferation.

537795-19-4
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

US 2002-176235

557795-19-4
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(SU 11248; composition and uses of galectin antagonists to augment

treatment of cancer or other proliferative disorders)

ANSWER 6 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) cell surface membrane receptors in cells of the specimen to such status, including presence or absence of a pre-cancerous state, presence or absence of a cancerous state, prognosis of a cancer, or responsiveness to treatment. Preferably, methods of the invention are implemented by using sets of binding compds. having releasable mol. tags that are specific for multiple components of one or more types of receptor dimers. After binding, mol. tags are release and sepd. from the assay mixt. for anal. S57795-19-4, SUI1248

Double bond geometry as shown.

ANSWER 7 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
557795-19-4 CAPLUS
H-Pytrole-3-carboxamide, N-{2-(diethylamino)ethyl}-5-{(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

Page 10 SAEED

## 10081147

L4 ANSWER 8 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2004:878170 CAPLUS

DOCUMENT NUMBER: 141:366237

Preparation of indolinone compounds for treatment of excessive osteolysis

NUTARY, Lesley: O'Farrell, Anne-Marie; Abrams, Tinya

SOURCE: USX/CO

DOCUMENT TYPE: Patent

EARGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INDORMATION: 1

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

|    | PA: | TENT I | NO.  |     |     | KIN | D   | DATE |      |     | APPL | ICAT | ION   | NO.  |     | D.  | ATE  |     |
|----|-----|--------|------|-----|-----|-----|-----|------|------|-----|------|------|-------|------|-----|-----|------|-----|
|    |     |        |      |     |     |     | -   |      |      |     |      |      |       |      |     | -   |      |     |
|    | US  | 2004   | 2099 | 37  |     | A1  |     | 2004 | 1021 | 1   | US 2 | 004- | 7809  | 17   |     | 2   | 0040 | 219 |
|    | WO  | 2004   | 0757 | 75  |     | A2  |     | 2004 | 0910 | 1   | WO 2 | 004~ | US 40 | 5283 |     | 2   | 0040 | 223 |
|    |     | W;     | ΑE,  | AG, | AL, | AM, | AT, | AU,  | AZ,  | BA, | вв,  | BG,  | BR,   | BW,  | BY, | BZ, | CA,  | CH, |
|    |     |        | CN,  | co, | CR, | CU, | CZ, | DE,  | DK,  | DM, | DZ,  | EC,  | EE,   | EG,  | ES, | FI, | GB,  | GD, |
|    |     |        | GE,  | GH, | GM, | HR, | ΗU, | ID,  | IL,  | IN, | IS,  | JP,  | ΚE,   | KG,  | KP, | KR, | ΚZ,  | LC, |
|    |     |        | LK,  | LR, | LS, | LT, | LU, | LV,  | ΜA,  | MD, | MG,  | MK,  | MN,   | MW,  | ΜX, | ΜZ, | ΝA,  | NI, |
|    |     |        | NO,  | NZ, | OM, | ₽G, | PH, | PL,  | PT,  | RO, | RU,  | sc,  | SD,   | SE,  | SG, | SK, | SL,  | SY, |
|    |     |        | ТJ,  | TM, | TN, | TR, | TT, | TZ,  | UΑ,  | UG, | US,  | υz,  | UG,   | vc,  | ٧N, | YU, | ZA,  | ZM, |
| ZW |     |        |      |     |     |     |     |      |      |     |      |      |       |      |     |     |      |     |
|    |     | RW:    | BW,  | GH, | GΜ, | KE, | LS, | MW,  | MZ,  | SD, | SL,  | SZ,  | TZ,   | υG,  | ZM, | ZW, | AM,  | ΑZ, |
|    |     |        | BY,  | KG, | ΚZ, | MD, | RU, | TJ,  | TM,  | AT, | BE,  | BG,  | CH,   | CY,  | CZ, | DE, | DK,  | EE, |
|    |     |        | ES,  | FI, | FR, | GB, | GR, | HU,  | IE,  | IT, | LU,  | MC,  | NL,   | PT,  | RO, | SE, | SI,  | sĸ, |
|    |     |        | TR,  | BF, | BJ, | CF, | CG, | CI,  | CM,  | GΑ, | GN,  | GQ,  | GW,   | ML,  | MR, | NE, | SN,  | TD, |
|    |     |        | TG,  | BF, | ВJ, | CF, | CG, | CI,  | CM,  | GΑ, | GN,  | GQ,  | G₩,   | ML,  | MR, | NE, | SN,  | TD, |
| TG |     |        |      |     |     |     |     |      |      |     |      |      |       |      |     |     |      |     |

PRIORITY APPLN. INFO.:

US 2003-448861P P 20030224

US 2004-780917 A 20040219

OTHER SOURCE(S):

MARPAT 141:366237

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB Disclosed is a method for treating excessive osteolysis in a patient, comprising administering to said patient an effective amount of a compound of

ound of formula (I) [wherein R = H, OH, alkyl, aryl, cycloalkyl, heteroaryl, alkoxy, heterocyclyl, amino; R1 = alkyl, halo, aryl, alkoxy, haloalkyl, haloalkoxy, cycloalkyl, heteroaryl, heterocyclyl, HO, COR8, NR9R10, NR9COR12, CONR9R10; R2 = alkyl, aryl, heteroaryl, COR8, SOR3;' (wherein R" = alkyl, aryl, heteroaryl, NR9R10, alkoxyl; R5 = H, alkyl, aryl, haloalkyl, cycloalkyl, heteroaryl, heterocyclyl, HO, COR8, (CHR)rR11; X = O, S; p, r = 0-3; q = 0-2; wherein R8 = OH, alkyl, aryl, heteroaryl, alkoxy, cycloalkyl, heterocyclyl; R9, R10 = H, alkyl, aryl, aminoalkyl, heteroaryl, cycloalkyl, heterocyclyl; R9, R10 = H, alkyl, aryl, aminoalkyl, heteroaryl, cycloalkyl, heterocyclyl; CNR9R10 together forms a ring consisting of the ring atoms selected from the group consisting of C, N,

ANSWER 8 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) (Therapeutic use); BIOL (Biological study); FREP (Preparation); USES (Uses)

(Uses)

(prepn. of indolinone compds. for treatment of excessive osteolysis, inhibiting phosphorylation of colony-stimulating factor-1 receptor (CSFIR), and treating cancers that express CSFIR)

RN 356068-94-5 CAPIUS

CN 1H-Pyrrole-3-carboxamide, 5-{(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME) NAME

Double bond geometry as shown

356068-97-8 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(ethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dlhydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

nd geometry as shown.

452104-42-6 CAPLUS
1H-Pyrrole-3-carboxamide, N-{3-{diethylamino}-2-hydroxypropyl]-5-{(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-{9CI}(CA\_INDEX\_NAME)

ANSWER 8 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
O, and S; R11 = OH, NH2, mono- or disubstituted amino, alkyl, aryl,
heteroaryl, alkoxy, cycloalkyl, heterocyclyl; R12 = alkyl, aryl,
heteroaryl, alkoxy, cycloalkyl, heterocyclyl; Z = OH, O-alkyl, NR3R4;
wherein R3, R4 = H, alkyl, aryl, heteroaryl, cycloalkyl, heterocyclyl; or
NR3R4 forms a ring consisting of the ring atoms selected from the group
consisting of CH2, N, O, and S, or Q1; wherein Y = CH2, O, N, S; Q = C,

n = 0-4; m = 0-3] or salts thereof. These compds. are useful for

n = 0-4; m = 0-3] or salts thereof. These compds. are useful for treating
excessive osteolysis, by inhibiting M-CSF mediated osteoclast development.

They are useful for inhibiting phosphorylation of colony-stimulating factor-1 receptor (CSF1R), and for treating cancers that express CSF1R. Thus, in a study on bone metastasis of cancer, 5-[5-Fluoro-2-oxo-1,2-dihydroindol-3-ylidenemethyl]-2,4-dimethyl-1H-pyrole-3-carboxylic acid (2-diethylaminoethyl)amide (II) at 80 or 40 mg/kg per day for 21 days inhibited the growth of 435/MAL-luc breast cancer cells in bone by 89% in mice in 41 days after inoculation with cancer cells. Formulations, e.g. hard gelatin capsule contg. II, were described.

IT 35008-94-59 356068-97-8P, 5-[([32]-5-Fluoro-2-oxo-1,2-dihydroindol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-ethylaminoethyl)amide 452104-42-6P, 5-[(5-Fluoro-2-oxo-1,2-dihydroindol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid N-(3-diethylamino-2-hydroxypropyl)amide 452104-85-7P,

5-[((32)-5-Fluoro-2-oxo-1,2-dihydroindol-3-ylidene)methyl]-2,4-dimethyl-1Hpyrrole-3-carboxylic acid [2-hydroxy-3-(morpholin-4-yl)propyl]amide
452104-86-89, 2,4-Dimethyl-5-((32)-2-oxo-1,2-dihydroindol-3ylidene)methyl]-H-pyrrole-3-carboxylic acid (2-hydroxy-3-(morpholin-4yl)propyl]amide 452104-87-99, 5-(((32)-5-Chloro-2-oxo-1,2dihydroindol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
N-[2-hydroxy-3-(morpholin-4-yl)propyl]amide 452104-88-09,

5-[((3Z)-5-Bromo-2-oxo-1,2-dihydroindol-3-ylidene)methyl]-2,4-dimethyl-1Hpyrrole-3-carboxylic acid N-[2-hydroxy-3-(morpholin-4-y1)propyl]amide 452104-89-1P, 2, 4-Dimethyl-5-(((32)-2-oxo-1,2-dihydroindol-3-ylidene)methyl]-1H-pyrrole-3-carboxylic acid N-[2-hydroxy-3-([1,2,3]triazol-1-yl)propyl]amide 452104-90-4P,

(1,2,3)triazol-1-yl)propyl]amide 452104-90-69,

5-{(32)-5-Fluoro-2-oxo-1,2-dihydroindol-3-ylidene)methyl]-2,4-dimethyl-1Hpyprole-3-carboxylic acid N-[2-hydroxy-3-(1H-[1,2,3]triazol-1yl)propyl]amide 452104-91-59, 5-[((32)-5-Chloro-2-oxo-1,2dihydroindol-3-ylidene)methyl]-2,4-dimethyl-1H-pyprole-3-carboxylic acid
N-[2-hydroxy-3-(1H-[1,2,3]triazol-1-yl)propyl]amide 452104-92-69
, 5-[((32)-5-Bromo-2-oxo-1,2-dihydroindol-3-ylidene)methyl]-2,4-dimethyl1H-pyprole-3-carboxylic acid N-[2-hydroxy-3-(1H-[1,2,3]triazol-1yl)propyl]amide 452105-23-69, 5-[((32)-5-Fluoro-2-oxo-1,2dihydroindol-3-ylidene)methyl]-2,4-dimethyl-1H-pyprole-3-carboxylic acid
N-[(25)-2-hydroxy-3-(morpholin-4-yl)propyl]amide 452105-24-79
499220-14-39 557795-19-49, 5-(5-Fluoro-2-oxo-1,2dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-pyprole-3-carboxylic acid
N-[(2-diethylaminoethyl) amide 587879-12-79, (2)-5-[(5-Fluoro-2-oxo-1,2dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-pyprole-3-carboxylic acid N-[2-(morpholin-4-yl) ethyl]amide
RL: FAC (Pharmacological activity); SPN (Synthetic preparation); THU

ANSWER 8 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-85-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-fluoro-1,2-dihydro-2-exo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI)(CA INDEX NAME)

Double bond geometry as shown

452104-86-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[{Z}]-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(CA INDEX NAME)

Double bond geometry as shown.

452104-87-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{Z}-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

ANSWER 8 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-88-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-{2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-89-1 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-{2-hydroxy-3-(HH-1,2,3-triazol-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-90-4 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-(2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 8 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452105-24-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[(ZR)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown

RN 499220-14-3 CAPLUS
CN Butanedioic acid, hydroxy-, (25)-, compd. with
N-[2-(diethylamino)ethyl]-5[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1Hpyrrole-3-carboxamide (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 342641-94-5 CMF C22 H27 F N4 O2

Absolute stereochemistry. Rotation (-).

L4 ANSWER 8 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-91-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-{2-hydroxy-3-{1H-1,2,3-triazol-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-92-6 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-{2-hydroxy-3-(H-1,2,3-triazol-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

452105-23-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[(2S)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

L4 ANSWER 8 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

557795-19-4 CAPLUS 1H-Pytrole-3-carboxamide, N-[2-{diethylamino}ethyl]-5-[{Z}-{5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene}methyl]-2,4-dimethyl- {9CI} (CA INDEX NAME)

Double bond geometry as shown.

587879-12-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-{2-(4-morpholinyl)ethyl}- (9CI) (CA INDEX

## 10081147

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| PATENT NO.             | KIND | DATE     | APPLICATION NO.   | DATE     |
|------------------------|------|----------|-------------------|----------|
|                        |      |          |                   |          |
| US 2004204407          | A1   | 20041014 | US 2004-793952    | 20040308 |
| PRIORITY APPLN. INFO.: |      |          | US 2003-452552P P | 20030307 |

OTHER SOURCE(S): MARPAT 141:350032

AB The title compds. [I; R1 and R2 combine to form (un)substituted optionally fused heterocyclic ring; R3-R5 = H, alkyl, hydroxyalkyl, etc.; or R3 and R4 may combine to form a cyclic 6-membered alicyclic ring which may be

ANSWER 9 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) y1) sulfony1]-1,2-dihydro-2-oxo-3H-indol-3-y1idene|methy1]-2,4-dimethy1-N-[3-(4-methy1-1-piperaziny1)propy1]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

PAGE 1-B

775321-60-3 CAPLUS

Double bond geometry as shown

775321-68-1 CAPLUS

//321-80-1 CAPUS Ha-Pyrrole-3-acetamide, 5-[(Z)-[5-[(2,3-dihydro-1H-indol-1-yl)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl|ethyl]- (9C1) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 9 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) substituted with one or more lower alkyl) that modulate the activity of protein kinases ("PKs") and are therefore useful in treating disorders related to abnormal PK activity (no biol. data), were prepd. General method of synthesis of the compds. I by condensation of oxindoles and aldehydes (prepn. of intermediates is given) is described. Eighty-two compds. I, e.g., II) were prepd. Pharmaceutical compns.comprising the compds. I, methods of treating diseases utilizing pharmaceutical compnising the comprising these compds. and methods of prepg. them are also disclosed. 775321-89-97 775321-59-09 775321-03-29 775321-73-99 775321-73-99 775321-73-99 775321-73-09-775321-73-09-775321-73-09-775321-73-09-775321-1

(Uses)
(preparation of 5-sulfonamido-substituted indolinone compds. as protein

kin selection in the selection of t

Double bond geometry as shown.

PAGE 1-B

775321-59-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-[5-[(5-bromo-2,3-dihydro-1H-indol-1-

ANSWER 9 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

775321-69-2 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl)-5-[(Z)-[5-[(Z)-3-dihydro-5-methoxy-1H-indol-1-yl)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

PAGE 1-B

PAGE 1-A

-NEt2

RN 775321-70-5 CAPLUS
CN 1H-Pyrrole-3-acetamide,
N-[2-[diethylamino]ethyl]-5-[(2)-[5-[(2,3-dihydro1H-indol-1-yl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 9 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued

RN 775321-73-8 CAPLUS
CN 1H-Pyrrole-3-acetamide,
5-[(2]-[5-[(3-chlorophenyl)methylamino]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 775321-75-0 CAPLUS
CN 1H-Pyrrole-3-acetamide, 5-{(2)-[5-{(2,3-dihydro-1H-indol-1-y1)sulfonyl}1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-{2-{4morpholinyl}-thyl}- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 775321-77-2 CAPLUS CN 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[{2}-[5-[{2,3-

L4 ANSWER 9 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 775321-81-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(Z)-[5-[(Z,3-dihydro-1H-indol-1-y1)sulfony1]-

1,2-dihydro-2-oxo-3H-indol-3-ylidene}methyl]-2,4-dimethyl-N-[3-(4-methyl-1-piperazinyl)propyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

PAGE 1-A

PAGE 1-B

RN 775322-01-5 CAPLUS

IH-Pyrrole-3-carboxamide, N-{2-(diethylamino)ethyl]-5-[(Z)-[5-[(5-fluoro-2,3-dihydro-1H-indol-1-yl)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 9 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) dihydro-1H-indol-1-yl)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 775321-78-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-(5-[(2,3-dihydro-1H-indol-1-yl)sulfonyl)1,2-dihydro-2-xo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 775321-80-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(Z)-(5-[(2,3-dihydro-1H-indol-1-yl)sulfonyl]1,2-dihydro-2-xo-3H-indol-3-ylidene]methyl]-2-ethyl-4-methyl-N-[2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 9 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 775322-06-0 CAPLUS
IH-Pyrrole-3-acetamide, 5-[(Z)-[5-(aminosulfonyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl)- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

RN 775322-07-1 CAPLUS

(N 1H-Pyrrole-3-acetamide, 5-[(2)-[1,2-dihydro-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]-(9CI) (CA INDEX NAME)

Double bond geometry as shown

RN 775322-08-2 CAPLUS
CN 1H-Pyrrole3-acetamide,
5-[(2)-[5-[(dimethylamino)sulfonyl]-1,2-dihydro-2oxo-3H-indol-3-ylidden|methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl](SCI) (CA INDEX NAME)

L4 ANSWER 9 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

RN 775322-12-8 CAPLUS
CN 1H-Pyrrole-3-acetamide,
5-((2)-(5-[((3-chlorophenyl)methylamino]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

775322-13-9 CAPLUS
1H-Pyrrole-3-acetamide, 5-[(2)-[5-(aminosulfonyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA

INDEX NAME)

Double bond geometry as shown.

ANSWER 9 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

775322-19-5 CAPLUS 1H-Pyrrole-3-acetamide, 5-[{Z}]-[1,2-dihydro-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]-(SCI) (CA INDEX NAME)

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Double bond geometry as sho

RN 775322-20-8 CAPLUS
CN 1H-Pyrrole-3-acetamide,
5-[(2]-[5-[(dimethylamino)sulfonyl]-1,2-dihydro-2oxo-3H-indol-3-ylidene[methyl]-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl](9CI) (CA INDEX NAME)

Double bond geometry as shown.

775322-25-3 CAPLUS 1H-Pyrrole-3-acetamide, N-[2-{diethylamino}ethyl]-5-[(2)-[5-[(dimethylamino)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-

ANSWER 9 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
775322-14-0 CAPLUS
1H-Pytrole-3-acetamide, N-(2-(diethylamino)ethyl)-5-[(Z)-[1,2-dihydro-5[(methylamino)aulfonyl]-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl(9CI) (CA INDEX NAME)

Double bond geometry as shown.

775322-15-1 CAPLUS
1H-Pyrrole-3-acetamide, 5-[(Z)-[5-(aminosulfonyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl}-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

RN 775322-18-4 CAPLUS
CN 1H-Pyrrole-3-acetamide,
5-[(Z)-[5-[[(3-chlorophenyl)methylamino]sulfonyl]-

1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl)-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 9 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN dimethyl- (9CI) (CA INDEX NAME) (Continued)

L4 ANSWER 10 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2004:740292 CAPLUS
DOCUMENT NUMBER: 141:265970
TITLE: Polymorphs of pyrrole-substituted 2-indolinone

INVENTOR (S):

kinase inhibitors
Sun, Changquan: Foster, Todd P.; Han, Fusen: Hawley,
Michael: Thamann, Tom
Pharmacia 6 Upjohn Company, USA
PCT Int. Appl., 36 pp.
CODEN: PIXXD2
Patent
Inglish
1

PATENT ASSIGNEE(S): SOURCE:

DOCUMENT TYPE: LANGUAGE:

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| PATE       | NT NO | ٥.   |     |     | KIN | D   | DATE |      |     |      |      |      |     |     |      |      |     |
|------------|-------|------|-----|-----|-----|-----|------|------|-----|------|------|------|-----|-----|------|------|-----|
|            |       |      |     |     |     | -   |      |      |     |      |      |      |     |     |      |      |     |
| WO 2       | 0040  | 764  | 10  |     | A2  |     | 2004 | 0910 |     | WO 2 | 004~ | US52 | 81  |     | 21   | 0040 | 223 |
|            | W: 3  | AΕ,  | AE, | AG. | AL. | AL, | AM.  | AM,  | AM, | AT.  | AT,  | AU,  | AZ. | AZ, | BA,  | BB,  | BG, |
|            |       |      |     |     |     |     | BY.  |      |     |      |      |      |     |     |      |      |     |
|            | Ċ     | cu.  | CU. | cz. | CZ. | DE. | DE.  | DK.  | DK. | DM.  | DZ,  | EC.  | EC. | EE. | EE.  | EG,  | ES. |
|            | 2     | ES.  | FI. | FI. | GB. | GD. | GE,  | GE.  | GH. | GM.  | HR.  | HR.  | HU. | HU, | ID.  | IL.  | IN. |
|            |       |      |     |     |     |     | KG,  |      |     |      |      |      |     |     |      |      |     |
|            |       |      |     |     |     |     | LU,  |      |     |      |      |      |     |     |      |      |     |
|            |       |      |     | NA. |     | ,   | ,    | ,    | ,   | ,    | ,    | ,    | ,   | ,   | ,    | ,    | ,   |
|            | RW: E |      |     |     |     | LS. | MW.  | MZ.  | SD. | SL.  | SZ.  | TZ.  | UG. | ZM. | ZW.  | AT.  | BE. |
|            |       |      |     |     |     |     | DK,  |      |     |      |      |      |     |     |      |      |     |
|            |       |      |     |     |     |     | SI,  |      |     |      |      |      |     |     |      |      |     |
|            |       |      |     |     |     |     | SN,  |      |     |      |      |      |     |     |      |      |     |
|            |       |      |     |     |     |     | SN,  |      |     | ٠.,  | υ,   | Ų.,  |     | U., | ٠,   | ٠,,, | ٠,  |
|            | 0042  |      |     |     |     |     |      |      |     |      | 004  | 1762 | 27  |     | 21   | 0040 | 712 |
|            |       |      |     |     | M.I |     | 2004 | 1223 |     |      |      |      |     |     |      |      |     |
| PRIORITY . | АРРЫ  | ١. ١ | NFO | . : |     |     |      |      |     | US 2 | 003- | 4488 | 63P |     | P 26 | 0030 | 224 |

US 2003-448863P P 20030224 US 2004-776337 A 20040212

The present invention relates to polymorphs of the 3-pyrrole-substituted 2-indolinone, 5-(5-fluoro-2-oxo-1,2-dihydro-indol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-pyrrolidin-1-ylethyl)amide (I). The phys. properties of polymorphs of I were determined by spectroscopic methods. methods. 753451-03-5 IT

RL: FMU (Formation, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); FORM (Formation, nonpreparative); USES

(Uses)
(polymorphs of pyrrole-substituted indolinone protein kinase inhibitors)
753451-03-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]-,monohydrochloride (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 11 OF 65
CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
DOCUMENT NUMBER:
11:23055
Semisolid oral formulations for immediate release containing indolylidenemethyl pyrrolepropionates
Martini, Alessandro; Ciocca, Cristina; Gatti, Paolo
Pharmacia Italia S.p.A., Italy
PCT Int. Appl., 31 pp.
CODEN: PIXXD2
DOCUMENT TYPE:
Pabet

Patent English 1

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO. APPLICATION NO. KIND DATE DATE WO 2004073592 WO 2004073592 W: AE, AI 20040130 PRIORITY APPLN. INFO.:

IT 2003-RM74

A 20030221

OTHER SOURCE(S):

MARPAT 141:230695

The present invention relates to a pharmaceutical composition suitable

oral administration, in the form of semisolid matrix, comprising an active ingredient poorly soluble in water and present in a quantity of from 15

45% by weight of the percent composition of the pharmaceutical composition; a

osition; a surfactant agent constituted by a polyglycolized glyceride; and a pharmaceutically acceptable hydrophilic carrier. A solid dispersion

L4 ANSWER 10 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

■ HC1

356068-94-5
RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); ΙT

(USES)

(polymorphs of pyrrole-substituted indolinone protein kinase inhibitors)

RN 356068-94-5 CAPLUS

CN 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX

Double bond geometry as shown.

ANSWER 11 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) on Gelucire 44/14 contg. 10% SU 6668 (I) was stable, homogeneous and had

dissoln. profile that assures that >90% of I is released within 45 min. 740153-03-5, SU 14813 Lmalate RE: PEP (Physical, engineering or chemical process); PRP (Properties); IT

PYP

{Physical process): THU (Therapeutic use): BIOL (Biological study): PROC (Process): USES (Uses)

(semisolid oral formulations for immediate release containing indolylidenmenthyl pyrrolepropionates)

RN 748153-83-5 CAPLUS

CN Butanedioic acid, hydroxy-, (2S)-, compd. with
5-[(5-fluoro-1,2-dihydro-2-

oxo-3H-indol-3-ylidene)methyl)-N-[(2S)-2-hydroxy-3-(4-morpholinyl)propyl]2,4-dimethyl-1H-pyrrole-3-carboxamide (9CI) (CA INDEX NAME)

CM 1

CRN 627908-92-3 CMF C23 H27 F N4 O4

Absolute stereochemistry. Double bond geometry unknown

2 CM

97-67-6 C4 H6 O5

Absolute stereochemistry. Rotation (-).

557795-19-4, SU 11248
RL: TRU (Therapeutic use): BIOL (Biological atudy); USES (Uses)
(semisolid oral formulations for immediate release containing
indolylidenemethyl pyrrolepropionates)
57795-19-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-{(Z)-(5-fluoro-1, 2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2, 4-dimethyl- (9CI) (CA INDEX

Double bond geometry as shown.

Page 16

SAEED

ANSWER 11 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

L4 ANSWER 12 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2004:565086 CAPLUS
DOCUMENT NUMBER: 141:123632
TITLE: Preparation of 3,5-Disubstituted-[1,2,4]-oxadiazoles and analogs as activators of caspases and inducers of apoptosis Cai, Sui Xiong; Zhang, Han-zhong; Kuemmerle, Jared INVENTOR(S): Zhang, Hong; Kemnitzer, William E. Cytovia, Inc., USA
PCT Int. Appl., 97 pp.
CODEN: PIXXD2
Patent
English
1 PATENT ASSIGNEE(S): SOURCE: DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. DATE APPLICATION NO. KIND DATE EMT NO. KIND DATE APPLICATION NO. DATE

2004058253 A1 20040715 W0 2003-US40308 20031218

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CC, CC, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MM, MW, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TM, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW

RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, WO 2004058253 US 2004127521 A1 20040701 US 2003-737865 US 2002-433953P 20031218 P 20021218 PRIORITY APPLN. INFO.: OTHER SOURCE(S): MARPAT 141:123632

II

Title compds. I (R1-3 = H, halo, haloalkyl, aryl, etc.; Q = S, O, amino; = heterocycle, carbocycle) are prepared For instance, 3-amino-4-

ANSWER 12 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) chlorobenzamidoxime (prepn. given) is reacted with 3-chlorothiophene-2-carbonyl chloride (pyridine, reflux, 50 min) to give II. II and other examples are potent caspase cascade activators and inducers of apoptosis in solid tumor cells, e.g., human breast cancer cell lines T-47D and

In Solid Cumbr Cells, 4.9., Number States Control States Care Stat

apoptosis)
557795-19-4 CAPLUS
HR-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl)-5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

Double bond geometry as shown.

ΙT

L4 ANSWER 13 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN ACCESSION NUMBER: 2004:534300 CAPLUS DOCUMENT NUMBER: 141:65034 TITLE: Substituted DOCUMENT NUMBER: 141:65094

TITLE: Substituted
1-benzoyl-3-cyano-pyrrolo(1,2-a)quinolines
and analogs as activators of caspases and inducers of apoptosis

INVENTOR(S): Cai, Sui xiong; Drewe, John A.; Jiang, Sungchun; Kasibhatla, Shailaja; Kuemmerle, Jared Daniel; Sirisoma, Nilantha Sudath; Zhang, Han-Zhong

PATENT ASSIGNEE(S): Cytovia, Inc., USA
SOURCE: PTXXD2

DOCUMENT TYPE: Patent
LANGUAGE: English

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

T

|      | PATENT   |       |      |     | KIN | D   | DATE |      |     | APPL | ICAT | ION   | NO. |     | D   | ATE  |     |
|------|----------|-------|------|-----|-----|-----|------|------|-----|------|------|-------|-----|-----|-----|------|-----|
|      |          |       |      |     |     | -   |      |      |     |      |      |       |     |     | -   |      |     |
|      | WO 2004  | 10551 | 63   |     | A2  |     | 2004 | 0701 |     | WO 2 | 003- | US 39 | 550 |     | 2   | 0031 | 212 |
|      | WO 2004  | 0551  | 63   |     | A3  |     | 2004 | 0826 |     |      |      |       |     |     |     |      |     |
|      | W:       | ΑE,   | AG,  | AL, | AM, | AT, | AU,  | AZ,  | BA, | BB,  | BG,  | BR,   | BW, | BY, | BZ. | CA.  | CH, |
|      |          | CN,   | co,  | CR, | CU, | CZ, | DE,  | DK,  | DM, | DZ.  | EC.  | EE.   | EG. | ES, | FI. | GB.  | GD. |
|      |          |       |      |     |     |     | ID,  |      |     |      |      |       |     |     |     |      |     |
|      |          |       |      |     |     |     | LV,  |      |     |      |      |       |     |     |     |      |     |
|      |          | NZ,   | OM,  | PG, | PH, | PL, | PT,  | RO,  | RU, | sc,  | SD,  | SE,   | SG, | SK, | SL. | SY.  | TJ. |
|      |          | TM,   | TN,  | TR, | TT, | TZ, | UA,  | UG,  | UZ, | VC,  | VN,  | YU,   | ZA, | ZM, | ZW  |      |     |
|      | RW:      | BW,   | GH,  | GM, | KE, | LS, | MW,  | MZ,  | SD, | SL.  | SZ,  | TZ,   | UG. | ZM. | ZW. | AM.  | AZ. |
|      |          |       |      |     |     |     | TJ,  |      |     |      |      |       |     |     |     |      |     |
|      |          |       |      |     |     |     | HU,  |      |     |      |      |       |     |     |     |      |     |
|      |          | TR,   | BF,  | ВJ, | CF, | CG, | CI,  | CM,  | GA, | GN,  | GQ,  | GW,   | ML, | MR. | NE. | SN.  | TD. |
| rG   |          |       |      |     |     |     |      |      |     |      |      |       |     | -   |     |      |     |
|      | US 2005  | 0147  | 59   |     | A1  |     | 2005 | 0120 |     | US 2 | 003- | 7332  | 29  |     | 2   | 0031 | 212 |
| PRIO | RITY APP | LN.   | INFO | .:  |     |     |      |      |     | US 2 | 002- | 4326  | 08P |     | P 2 | 0021 | 212 |
|      |          |       |      |     |     |     |      |      |     |      |      |       |     |     |     |      |     |

OTHER SOURCE(S): MARPAT 141:65094

BY The invention discloses substituted 1-benzoyl-3-cyanopyrrolo[1,2-a]quinolines and analogs thereof. Compds. of the invention are activators

of caspases and inducers of apoptosis. Therefore, the compds. of the invention can be used to induce cell death in a variety of clin. conditions in which uncontrolled growth and spread of abnormal cells occurs. Compound prepn is described.

IT 557795-19-4, SU 11248

RL: PAC (Pharmacological activity): THU (Therapeutic use): BIOL (Biological study): USES (Uses)

(benzoylcyanopyroloquinolines and analogs as activators of caspases and inducers of apoptosis)

RN 557795-19-4 CAPIUS

CN 1H-Pytrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[{Z}]-(5-fluoro-1, 2-dihyto-2-oxo-3H-indol-3-ylidene)methyl]-2, 4-dimethyl- (9CI) (CA INDEX NAME)

ANSWER 13 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

ANSWER 14 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
557795-19-4, SU 11248
RL: TRU (Therapeutic use); BIOL (Biological study); USES (Uses)
(KIT tyrosine kinase inhibitor; methods for assessing anti-cancer activity of KIT tyrosine kinase inhibitor, gastrointestinal stromal tumor treatment, and assessing cancer progression, using gene expression profiling)
557795-19-4 CAPLUS
HI-Pyrrole-3-carboxamide, N-(2-(diethylamino)ethyl)-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 14 OF 65
ACCESSION NUMBER:
DOCUMENT NUMBER:
141:33766
Hethods for assessing the anti-cancer activity of a KIT tyrosine kinase inhibitor, gastrointestinal atomal tumor treatment, and assessing cancer progression, using gene expression profiling Eisenberg, Burton: Von Mehren, Margaret; Frolov, Andrey; Godwin, Andrew
FOX Chase Cancer Center, USA
PATENT ASSIGNEE(S):
SOURCE:
DOCUMENT TYPE:
LANGUAGE:
PANTLY ACC. NUM. COUNT:
FAMILY DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE 2004045545 A2 i W: AE, AG, AL, AM, AT, CN, CO, CR, CU, CZ, GH, GM, HR, HU, ID, LR, IS, LT, LU, LV, OM, PG, PH, PL, PT, TN, TR, TT, TZ, UA, RW: BW, GH, GM, KE, LS, BY, KG, KZ, MD, RU, ES, FI, FR, GB, GR, TR, BF, BJ, CF, CG, 20040603 20040812 , AU, AZ, , DE, DK, , II., IN, , MA, MD, , RO, RU, , UG, US, , MW, MZ, , TJ, TM, , HU, IE, , CI, CM, WO 2004045545 WO 2004045545 WO 2003-US36820 20031118 BG, BR, BW, BY, B2, CA, CH, EC, EE, ES, FI, GB, GD, GE, KE, KG, KP, KR, KZ, LC, LK, NN, HW, MX, MZ, NI, NO, NZ, SE, SG, SK, SL, SY, TJ, TM, VN, YU, ZA, ZM, ZW, ZM, AB, AZ, BG, CH, CY, CZ, DE, DK, EE, MC, NL, PT, RO, SE, SI, SK, GQ, GW, ML, MR, NE, SN, TD, BA, BB, DM, DZ, IS, JP, MG, MK, SC, SD, UZ, VC, SD, SL, AT, BE, IT, LU, GA, GN, TG PRIORITY APPLN. INFO.: US 2002-427326P P 20021118 The present invention provides novel methods for the treatment of cancer, methods for screening compds. having anti-cancer activity, and methods of assessing cancer progression. In accordance with the present invention, method of assessing the anti-cancer activity of a KIT tyrosine kinase inhibitor in a biol. sample comprising a tumor cell is provided. In a preferred embodiment, the tumor is a gastrointestinal stromal tumor and the KIT tyrosine kinase inhibitor is imatinib, SU11248 (Sugen Pharmaceuticals), or a pharmaceutically acceptable salt thereof. DNA microarrays revealed 148 genes that were differentially expressed between untreated and imatinib-treated human GIST colls, in vitro. One of these genes, Sprouty4A (SPRY4A) a regulator of tyrosine kinase-mediated signaling pathways, was dramatically down-regulated. A biomarker MAFbx was up-regulated in response to imatinib treatment. In addition, which inhibited KIT phosphorylation without affecting the total level of KIT protein. The inventors proposed a method for determining the efficacy anticancer treatment comprising detection of an alteration in phosphorylation of a biomarker (such as decrease in GAB1 phosphorylation).

L4 ANSWER 15 OF 65
ACCESSION NUMBER:
DOCUMENT NUMBER:
11112:

INVENTOR(S):
ADFAMENT ASSIGNEE(S):
SOURCE:
DOCUMENT TYPE:
LANGUAGE:
LANGUAGE:
ADFAMENT ACC. NUM. COUNT:

DOCUMENT TYPE:
LANGUAGE:
BANGUAGE:
BANG

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO. DATE KIND APPLICATION NO. DATE W0 2004045523
W0 2004045523
W: AE, AG,
CN, CO,
GE, GH,
LK, LR,
NZ, OM,
TM, TM,
RW: BM, GH,
BY, KG,
ES, FI,
TR, BF, A2 A3 20040603 WO 2003-US36526 20031114 A3 20040930
AM, AT, AU, AZ, BA, BB,
CU, CZ, DE, DK, DM, DZ,
RR, HU, ID, IL, IN, IS,
LT, LU, LV, MA, MD, MG,
PH, PL, PT, RO, RU, SC,
TT, TZ, UA, UG, US, UZ,
KE, LS, MW, MZ, SD, SL,
MD, RU, TJ, TM, AT, BE,
GB, GR, HU, IE, IT, LU,
CF, CG, CI, CM, GA, GN, BG, BR, BW, EC, EE, EG, JP, KE, KG, MK, MN, MW, SD, SE, SG, VC, VN, YU, SZ, TZ, UG, BG, CH, CY, BZ, CA, CH, FI, GB, GD, KR, KZ, LC, MZ, NI, NO, SL, SY, TJ, ZM, ZW ZW, AM, AZ, DE, DK, EE, SE, SI, SK, AL, CR, GM, LS, PG, TR, GM, KZ, FR, BJ, BY, ES, KP, MX, SK, ZA, ZM, CZ, RO, MC, NL, PT, GQ, GW, ML, NL 1024779 NL 1024779 US 2004152759 A1 C2 20040518 NL 2003-1024779 20031114 20041109 A1 20040805 US 2003-712296 US 2002-426386P 20031114 P 20021115 PRIORITY APPLN. INFO.: OTHER SOURCE(S): MARPAT 141:1206

The invention relates to a method of treating cancer by administering a combination of an indolinone compound with another chemotherapeutic

Page 18 SAEED ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
The combination of an indolinone compd. I (R = H, OH, alkyl, aryl,
cycloalkyl, heteroaryl, alkoxy, heterocycle, amino: R1 = alkyl, halo,
alkoxy, etc.: R2 = alkyl, aryl, heteroaryl, etc.: R5 = H, alkyl, aryl,
haloalkyl, cycloalkyl, etc.: X = O, S: p = 0, 1, 2, 3; q = 0, 1, 2; Z =
OH, -O-alkyl, -NN3R4: R3, R4 = H, alkyl, aryl, heteroaryl, cycloalkyl,
heterocycle, or together with N form a ring) with another
otherapeutic
agent provides an enhanced effect in treating cancer patients. Mice
implanted with MC-1 human breast carcinoma fragments were treated wide
docetaxel and 5-(5-fluoro-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide (prepn.
given).

dimethyl-1H-pyrrole-3-carboxylic acid (2-01cm, given).

IT 627908-92-3F 674778-85-9P
RL: BSU (Biological study, unclassified); PAC (Pharmacological activity);
SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(as indolinone compound; cancer therapy using combination administration
of indolinone compuds, with chemotherapeutic agents for cell proliferation disorders)
RN 627908-92-3 CAPUUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl)-N-[(25)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(SCI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown

674778-85-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-M-[(2R)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN NAME)

RN 346405-32-1 CAPLUS
CN | H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-(2-(1-pyrrolidinyl)ethyl)- (9CI) (CINDEX | CONTROL | CONTROL

RN 356069-16-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(ethylamino) ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-94-5 342641-94-5D, acceptable salts, solvates, hydrates 346405-32-1 346405-32-1D, acceptable salts, solvates, hydrates 355069-16-4 355069-16-4D, acceptable salts, solvates, hydrates 452104-43-70, acceptable salts, solvates, hydrates 452104-43-70, acceptable salts, solvates, hydrates 515338-92-6D, acceptable salts, solvates, hydrates 515338-92-6D, acceptable salts, solvates, hydrates 627908-92-3D, acceptable salts, solvates, hydrates 637924-92-1

697224-02-1

RL: BSU Gological study, unclassified); PAC (Pharmacological activity);
THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(as indolinone compound; cancer therapy using combination
administration
of indolinone compds. with chemotherapeutic agents for cell
proliferation disorders)
RN 342641-94-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-((5-fluoro-1,2dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX
NAME)

342641-94-5 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 346405-32-1 CAPLUS
CN 1H-Fyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl-N-(2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX

ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356069-16-4 CAPLUS
1H-Pyrrole-3-carboxamide,
(ethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

452104-43-7 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl}-2,4-dimethyl-(9CI)(CA INDEX NAME)

452104-43-7 CAPLUS

Page 19

ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) lH-Pyrrole-3-carboxamide, 5-{(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene] methyl|-N-{2-hydroxy-3-(4-morpholinyl)propyl|-2,4-dimethyl-(9CI) (CA INDEX NAME)

452104-45-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI)
(CA INDEX NAME)

ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene)methyl]-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAME)

515138-82-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-{2-(4-morpholinyl)ethyl}- (9CI) (CA INDEX NAME)

627908-92-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-{(23)-2-hydroxy-3-(4-morpholinyl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

14 ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-45-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI)(CA INDEX NAME)

515138-82-6 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-

ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

674778-85-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[(2R)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown

697224-82-1 CAPLUS Butanedioic acid, hydroxy-, compd. with N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrole-3-carboxamide (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 342641-94-5 CMF C22 H27 F N4 O2

CRN 6915-15-7 CMF C4 H6 O5

ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

но2с-сн-сн2-со2н

356068-97-8P, 5-(5-Fluoro-2-oxo-1,2-dihydroindol-(3Z)ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
(2-ethylaminoethyl)amide 452104-62-68 452104-85-97
452104-69-69 452104-97-99 452104-88-09
452104-89-19 452104-90-69 452104-91-59
452104-89-19 452104-90-69 452104-91-59
452104-89-69 452103-25-69 452105-24-79
RL: SPN (Synthetic preparation); PREP (Preparation)
(cancer therapy using combination administration of indolinone compds.
with chemotherapeutic agents for cell proliferation disorders)
356068-97-8 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(ethylamino)ethyl]-5-{(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-42-6 CAPLUS 1H-Pyrrole-3-carboxamide, N-[3-(diethylamino)-2-hydroxypropyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-(9CI)(CA INDEX NAME)

452104-85-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

452104-89-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-((2)-(1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[2-hydroxy-3-(HH-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

452104-90-4 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

452104-91-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-((Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-86-8 CAPLUS
1H-Pyrole-3-carboxamide, 5-[{Z}]-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(CA INDEX NAME)

Double bond geometry as shown.

452104-87-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{Z}]-{5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

452104-88-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-92-6 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl}-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

452105-23-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{{Z}}-{5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-H-{{Z}}-2-hydroxy-3-{4-morpholinyl}propyl}-2,4-dimethyl-{GL} (AC INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

452105-24-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-N-[(2R)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

L4 ANSWER 15 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

ANSWER 16 OF 65 CAPLUS COPYRIGHT 2005 ACS ON STN SSION NUMBER: 2004:252326 CAPLUS MENT NUMBER: 140:276195 ACCESSION NUMBER:

DOCUMENT NUMBER: TITLE: 140:276195
Formulations comprising an indolinone compound
Gatti, Paolo
Pharmacia Italia S.P.A., Italy
PCT Int. Appl., 109 pp.
CODEN: PIXEND
Patent INVENTOR(S):

PATENT ASSIGNEE (S):

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION: English

| PA      | TENT  | NO.   |      |     | KIN | D   | DATE |      |     | APPL | ICAT | ION  | NO. |     | D    | ATE  |     |
|---------|-------|-------|------|-----|-----|-----|------|------|-----|------|------|------|-----|-----|------|------|-----|
| WO      | 2004  | 0241  | 27   |     | A2  | _   | 2004 | 0325 | ,   | WO 2 | 003- | 1852 | 93  |     | 21   | 0030 |     |
| WO      | 2004  | 0241  | 27   |     | A3  |     | 2004 | 0603 |     |      |      |      |     |     | _    |      |     |
|         | W:    | ΑE,   | AG,  | AL, | AM, | AT, | ΑU,  | ΑZ,  | BA, | BB,  | BG,  | BR,  | BY, | BZ, | CA,  | CH,  | CN, |
|         |       | co,   | CR,  | CU, | CZ, | DE, | DK,  | DM,  | DZ, | EC,  | EE,  | ES,  | FI, | GB, | GD,  | GΕ,  | GH, |
|         |       | GM,   | HR,  | ΗU, | ID, | IL, | IN,  | IS,  | JP, | ΚE,  | KG,  | KP,  | KR, | ΚZ, | LC,  | LK,  | LR, |
|         |       | LS,   | LT,  | LU, | LV, | MA, | MD,  | MG,  | MK, | MN,  | MW,  | MX,  | MZ, | NI, | NO,  | NZ,  | OM, |
|         |       | PG,   | PH,  | PL, | PT, | RO, | RU,  | sc,  | SD, | SE,  | SG,  | SK,  | SL, | SY, | TJ,  | TM,  | TN, |
|         |       | TR,   | TT,  | TZ, | UA, | UG, | US,  | UZ,  | VC, | VN,  | YU,  | ZA,  | ZM, | ZW  |      |      |     |
|         | RW:   | GH,   | GM,  | KE, | LS, | MW, | MZ,  | SD,  | SL, | SZ,  | TZ,  | UG,  | ZM, | ZW, | AM,  | AZ,  | BY, |
|         |       | KG,   | KZ,  | MD, | RU, | TJ, | TM,  | ΑT,  | BE, | BG,  | CH,  | CY,  | CZ, | DE, | DK,  | EE,  | ES, |
|         |       | FI,   | FR,  | GB, | GR, | ΗU, | IE,  | IT,  | LU, | MC,  | NL,  | PT,  | RO, | SE, | SI,  | SK,  | TR, |
|         |       | BF,   | ВJ,  | CF, | CG, | CI, | CM,  | GΑ,  | GN, | GQ,  | GW,  | ML,  | MR, | NE, | SN,  | TD,  | TG  |
| NL      | 1024  | 261   |      |     | A1  |     | 2004 | 0311 | 1   | NL 2 | 003- | 1024 | 261 |     | 24   | 0030 | 910 |
| US      | 2004  | 2299: | 30   |     | A1  |     | 2004 | 1118 | 1   | US 2 | 003- | 6588 | 01  |     | 21   | 0030 | 910 |
| PRIORIT | Y APP | LN.   | INFO | .:  |     |     |      |      | 1   | US 2 | 002- | 4211 | 33P | -   | P 20 | 0020 | 910 |

OTHER SOURCE(S):

MARPAT 140:276195

AB The present invention features formulations of indolinones suitable for parenteral or oral administration. The formulations and the compds. themselves are useful for the treatment of protein kinase related disorders. For example, 5-(5-fluoro-2-oxo-1,2-dihydro-1ndol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylamino)-amide (1) was wet granulated with mannitol, croscarmellose sodium and polyvinylpyrrolidone, the granules were dried, blended with magnesium stearate, and filled in capsules; each capsule contained 50 mg, 75 mg, or 200 mg I.

IT 34264-94-5 452104-85-7 452105-23-6 452105-24-7 499220-14-3 627908-92-3 674778-85-9

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (preparation of oral and parenteral compns. of indolinone compound)

RN 342641-94-5 CAPLUS

CN 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

ANSWER 16 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-85-7 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{{Z}}-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-{2-hydroxy-3-{4-morpholinyl}propyl}-2,4-dimethyl-(CA INDEX NAME)

Double bond geometry as shown

452105-23-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-((Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-((ZS)-2-hydroxy-3-(4-morpholinyl)propyl)-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown

452105-24-7 CAPLUS 1H-Pyrrole-3-carboxamide, 5-([Z])-{5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[[ZR])-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

ANSWER 16 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

499220-14-3 CAPLUS
Butanedioic acid, hydroxy-, (2S)-, compd. with
-(diethylamino)ethyl]-5[(5-fluor-0,1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1Hpyrrole-3-carboxamide (1:1) (9CI) (CA INDEX NAME)

CRN 342641-94-5 CMF C22 H27 F N4 O2

CM 2

Absolute stereochemistry. Rotation (-).

627908-92-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-N-{(2S)-2-hydroxy-3-(4-morpholinyl)propyl}-2,4-dimethyl-(SCI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

10081147

ANSWER 16 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

674778-85-9 CAPLUS
1R-Pyrrole-3-carboxamide, 5-{(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-((2R)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(SCI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

ANSWER 17 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

REFERENCE COUNT:

THERE ARE 422 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE REFORMAT

ANSWER 17 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN SSION NUMBER: 2004:197494 CAPLUS ACCESSION NUMBER:

ACCESSION NUMBER: 2004:197494 CAPJUS

DOCUMENT NUMBER: 141:235330

TITLE: Emerging roles of targeted small molecule protein-tyrosine kinase inhibitors in cancer therapy AUTHOR(S): Smith, John K., Mamoon, Naila H.; Duhe, Rey J. CORPORATE SOURCE: Department of Pharmacology and Toxicology, University of Hississippi Medical Center, Jackson, MS, 39216-4505, USA

SOURCE: Oncology Research (2003), 14(4/5), 175-225

COODEN: ONRES! ISSN: 0965-0407

PUBLISHER: Cognizant Communication Corp.

DOCUMENT TYPE: Journal; General Review LandungGE: English

AB A review. Targeted protein-tyrosine kinase inhibitors (PTKIs) comprise a new, rapidly evolving class of low mol. weight anticancer drugs. Two members

of this class, imatinib (Gleevec) and gefitinib (tresss), are currently approved for market use in the United States. This review discusses the scientific history behind these two PTKI drugs, including the role of the targeted kinase in cancer eticl., the biochem. of selective inhibition, the evaluation of clin. efficacy, and the mechanisms whereby drug resistance has emerged. Other PTKIs undergoing clin. evaluation are also described, including epidermal growth factor receptor kinase inhibitors (eriotinib, PKI166, and CI-1033) and PTKIs designed to disrupt tumor vacularization (SUS161, SUS668, SU1248, PTK707, and 2D6474). How might one apply current knowledge to the efficient development of new agents that would target as-yet-unexploited oncogenic PTKs such as chimeric anaplastic leukemia kinases or Janus kinases. Ideally, the targets should contain structurally distinct drug interaction epitopes, although it is not necessary that these epitones he under the protect of th

dd contain structurally distinct drug interaction epitopes, although it is not necessary that these epitopes be unique to a single target, because effective drugs may inhibit multiple kinases involved in an oncogenic process. Oral availability is a highly desirable feature because daily oral administration can maintain a sustained efficacious plasma

entration, whereas intermittent parenteral administration may not. Perhaps most importantly, one must verify the presence of an appropriate mol. target on

a case-by-case basis before selecting a patient for PTKI therapy. Thus, the development of molecularly targeted diagnostic tools will be crucial to the ultimate success of molecularly targeted PTKI therapy. 57799-19-4, SU 11248
RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (SU 11248; epidermal growth factor receptor kinase inhibitor CI-1033 IT

is designed to disrupt tumor vascularization SU11248 and used in treatment

tment
of cancer therapy)
557795-19-4 CAPLUS
HI-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

Double bond geometry as shown.

L4 ANSWER 18 OF 65 CAPLUS COPYRIGHT 2005 ACS ON STN ACCESSION NUMBER: 2004:182368 CAPLUS COPYRIGHT 2005 ACS ON STN 1004:182368 CAPLUS 140:229401 THILE: 1igand-binding

Three hybrid assay system for isolating

polypeptides and for isolating small mol. ligands Come, Jon H.; Becker, Frank; Kley, Nikolai A.; Reichel, Christoph

VSA
U.S. Pat. Appl. Publ., 238 pp., Cont.-in-part of U.S.
Ser. No. 31,177.
CODEN: USXXCO
Patent PATENT ASSIGNEE(S): SOURCE:

DOCUMENT TYPE:

English 5

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| PATENT NO.                              | KIND | DATE     | APPLICATION NO. | DATE        |
|---|------|----------|-----------------|-------------|
| US 2004043388<br>US 2003165873          |      | 20040304 |                 |             |
| US 2004266854<br>PRIORITY APPLN. INFO.: | Al   | 20041230 | US 2004-820453  |             |
|   |      |          | US 2001-278233P | P 20010323  |
|   |      |          | US 2001-329437P | P 20011015  |
|   |      |          | US 2002-91177   | A2 20020304 |
|   |      |          | US 2001-336962P | P 20011203  |
|   |      |          | WO 2002-US6677  | A2 20020304 |
|   |      |          | US 2002-234985  | A2 20020903 |
|   |      |          | WO 2002-US33052 | A2 20021015 |
|   |      |          | US 2003-460921P | P 20030407  |
|   |      |          | US 2003-531872P | P 20031223  |

The invention provides compns. and methods for isolating ligand-binding polypeptides for a user-specified ligand, and for isolating small molligands for a user-specified target polypeptide using an improved class

hybrid ligand compds. Preparation of compds., e.g a methotrexate moiety linked

ed
by a polyethylene gycol moiety to dexamethasone, is described.
295799-47-2D, conjugates
RL: BUU (Biological use, unclassified); THU (Therapeutic use); BIOL
(Biological study); USES (Uses)
(three hybrid assay system for isolating ligand-binding polypeptides
and for isolating small mol. ligands)
295799-47-2 CAPLUS

Z95799-47-2 CAPLUS
ZH-Indol-2-one, 3-[(4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2yl]methylene]-1,3-dihydro-6-(3-pyridinyl)- (9CI) (CA INDEX NAME)

Page 23 SAEED

#### 10081147

ANSWER 18 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

ANSWER 19 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

REFERENCE COUNT: THIS

THERE ARE 17 CITED REFERENCES AVAILABLE FOR

FORMAT

RECORD. ALL CITATIONS AVAILABLE IN THE RE

ANSWER 19 OF 65 CAPLUS COPYRIGHT 2005 ACS ON STN SSION NUMBER: 2004:172077 CAPLUS MENT NUMBER: 140:357149

ACCESSION NUMBER: DOCUMENT NUMBER:

140:357149
Amidations Using N,N'-Carbonyldiimidazole: Remarkable Rate Enhancement by Carbon Dioxide
Rate Enhancement by Carbon Dioxide
Vaidyanathan, Rajappa; Kalthod, Vikram G.; Ngo, Duc P.; Manley, Jerad M.; Lapekas, Sean P.
Chemical Research and Development, Pfizer Inc.,
Kalamazoo, HI, 49001, USA
Journal of Organic Chemistry (2004), 69(7), 2565-2568
CODEN: JOCEAH; ISSN: 0022-3263
American Chemical Society
Journal
English TITLE:

AUTHOR (S):

CORPORATE SOURCE:

SOUTHER.

PUBLISHER

DOCUMENT LANGUAGE:

ΑB Carbon dioxide catalyzes the reaction of imidazolides with amines to form amides. A substantial rate enhancement is observed in the presence of

compared to the CO2-free case. The scope and limitations of this reaction

Compared to the CO2-free case. The scope and limitations of this reaction
are discussed. For example, amidation of
5-formyl-2,4-dimethyl-1H-pyrrole3-carboxylic acid (I) in the presence of 1,1'-carbonylbis[1H-imidazole] and carbon dioxide gave N-[2-(diethylamino)ethyl]-5-[[[2-(diethylamino)ethyl]-5,4-dimethyl-1H-pyrrole-3-carboxamide (II). Hydrolysis of II gave N-[2-(diethylamino)ethyl]-2,4-dimethyl-1H-pyrrole-3-carboxamide. Reaction of II with
5-fluoro-1,3-dihydro-2H-indol-2-one gave N-[2-(diethylamino)ethyl]-5-[[5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-1H-pyrrole-3-carboxamide. The effect of 1-[(2,4-diethyl-1H-pyrrol-3-yl)carbonyl]-1H-imidazole on the amidation kinetics was evaluated.
IT 34261-94-5P, N-[2-(Diethylamino)ethyl]-5-[(5-fluor-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-1H-pyrrole-3-carboxamide
RL: SPN (Synthetic preparation); PREP (Preparation)
presence of carboxamides from carboxylic acids and amines in presence of carbonylbis[imidazole] and carbon dioxide as catalyst)

ence of carbonylbis[imidazole] and carbon dioxide as catalyst)
342641-94-5 CAPLUS
HI-Pyrcole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

L4 ANSWER 20 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN ACCESSION NUMBER: 2004:162557 CAPLUS COPYRIGHT 2005 ACS on STN 2004:16257 CAPLUS COPYRIGHT 2005 ACS ON STN 2004:1 140:195469
Phosphatidylinositol 3-kinase antagonists as radiosensitizers
Hallahan, Dennis E.; Tan, Jiahuai
Vanderbilt University, USA
PCT Int. Appl., 77 pp.
CODEN: PIXXD2
Patent
English
1

INVENTOR (S)

PATENT ASSIGNEE(S): SOURCE:

DOCUMENT TYPE:

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE MO 2004016211 A2 20040226 WO 2003-USZ5015 20030808 WO 2004016211 A3 20040715 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MK, MX, MX, NX, NO, NZ, OM, PG, PH, PI, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, VU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NI, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG PRIORITY APPLN. INFO.:

A method for increasing radiosensitivity of a target tissue in a subject via administration of a phosphatidylinositol 3-kinase (PI3K) antagonist

a target tissue in a subject. Also provided are methods for suppressing tumor growth and for inhibiting tumor blood vessel growth via administration of a PI3K antagonist. 557795-19-4, SU11248

557795-19-4, SU11248
RE: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(PI3K antagonists as radiosensitizers targeted to tumor vascular
endothelium)
557795-19-4 CAPLUS
H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX
NAME)

L4 ANSWER 21 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2004:157243 CAPLUS
DOCUMENT NUMBER: 141:16612
Gene expression profiling of human colon xenograft tumors following treatment with SU11248, a multitargeted tyrosine kinase inhibitor
AUTHOR(S): Morito, Alysas M.; Tan, Nguyen; West, Kristina; McArthur, Grant; Toner, Guy C.; Manning, William C.; Smolich, Bewerlp D.; Cherrington, Julie M.
CORPORATE SOURCE: Department of Preclinical Research and Exploratory Development, SUGEN, Inc., South San Francisco, CA, 94080, USA
SOURCE: Oncogene (2004), 23(8), 1618-1626
CODEN: ONCMES; ISSN: 0950-9232
PUBLISHER: Nature Publishing Group
DOCUMENT TYPE: Journal
LANGUAGE: English
AB Biomarkers that indicate biol. activity and/or efficacy are a potentially useful tool in the development of molecularly targeted therapeutics. It is useful, though challenging, to identify biomarkers during preclin. development in order to impact decision-making during early clin. development. SU11248 is an oral, selective multitargeted tyrosine kinase inhibitor currently in Phase II oncol. clin. trials. It exhibits direct antitumor and antiangiogenic activity via inhibition of the receptor tyrosine kinases PDGFR, VEGFR, KIT and FLF3. To identify clin. translatable biomarkers of SU11248 activity, expression profiling was performed on Colo205 human xenograft tumors following treatment with SU11248. Over 100 transcripts changed in abundance in SU11248 s

ared
to vehicle-treated tumors. Nine candidate transcripts, chosen based on
putative function, were also analyzed and validated by TagMan. One such
potential biomarker, cadherin-ll, was further evaluated at the protein
level and was found to have increased expression in xenograft tumors

level and was found to have increased expression in xenograft tumors in SU1248 treatment. Interestingly, cadherin-II expression was also detected via immunohistochem. snal. of archived solid tumors, indicating the tech. feasibility of translating this putative biomarker to clin. studies. Importantly, SU11248 treatment also resulted in increased expression of cadherin-II protein in human tumor biopsies in three out of seven patients examined and confirms the feasibility of using transcriptional profiling of preclin. models to identify clin. translatable biomarkers.

557795-19-4, SU11248
RI: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(gene expression profiling of human colon xenograft tumors following treatment with SU11248, a multitargeted tyrosine kinase inhibitor)

557795-19-4 CAPLUS
IH-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dibydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NOME)

Double bond geometry as shown.

L4 ANSWER 22 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2004:143270 CAPLUS
DOCUMENT NUMBER: 140:197593

PDGFRa oncokinase fusion protein associated with hyperproliferative disease and as imatinib mesylate target in EOL-1 cell
RINVENTOR(S): Briesewitz, Roger; Griffin, John H.
PATENT ASSIGNEE(S): Theravance, Inc., USA
SOURCE: POT Int. Appl., 95 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: PRANLLY ACC. NUM. COUNT: 1

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

APPLICATION NO. PATENT NO. KIND DATE PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2004015082 A2 20040219 WO 2003-US24992 20030808

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, CM, CM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, LM, PT, RO, SE, SI, SK, ST, US, CM, BF, BJ, CF, CG, CI, CM, GA, GM, GQ, GW, ML, MR, NE, SN, TD, TG

US 2004045044 A1 20040304 US 2002-402330P F 20020809

FRIORITY APPLN. INFO: US 2003-440491P

Oncokinase fusion protein associated with hyperproliferative disorders

provided. The fusion polypeptides have a C-terminal tyrosine kinase domain fused to an N-terminal domain that is not normally fused to the C-terminal tyrosine kinase domain and they possess constitutively activated tyrosine kinase activity. The invention provides sequence of protein NM\_030917 fused with platelet-derived growth factor receptor a from human. The invention also identified deletion of 1 megabase fuses NM\_030917 and exon 12 of PDGFRG on human chromosome 4. Also provided are methods of diagnosing disease conditions associated with the fusion polypeptides. In addition, screening assays for identifying

useful for treating disease conditions associated with such fusion polypeptides and polynucleotides are provided. Furthermore, methods of treating disease conditions associated with the presence of the fusion polypeptides are provided.

537795-19-4, SU11248

RL: BSU [Bological study, unclassified]; BIOL (Biological study) (reducing activity of fusion protein by; PDGFRe oncokinase fusion protein associated with hyperproliferative disease and as imatinib mesylate target in EOL-1 cell)

557795-19-4 CAPLUS

1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 21 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

REFERENCE COUNT:

50 THERE ARE 50 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

ANSWER 22 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) -NEt2

Page 25 SAEED

## 10081147

L4 ANSWER 23 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
DOCUMENT NUMBER:
110:163705
Process for preparation of isotopically labeled indolinone derivatives
GITIONE, Danilo; Pignatti, Alberto; Fontana, Erminia
PATENT ASSIGNEE(S):
PATENT ASSIGNEE(S):
PATENT ASSIGNEE(S):
PATENT ASSIGNEE (S):
PARTIN TYPE:
PATENT NUMBER:
PARTIN TYPE:
PATENT NUMBERATION:
PATENT NUMBERATION:

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

|          |               |     |      |     |      |      |      |      |     | •    |      |      |     |     |          |      |     |  |
|----------|---------------|-----|------|-----|------|------|------|------|-----|------|------|------|-----|-----|----------|------|-----|--|
| PAT      | KIND DATE     |     |      |     | APPL | DATE |      |      |     |      |      |      |     |     |          |      |     |  |
|          |               |     |      |     |      | -    |      |      |     |      |      |      |     |     | -        |      |     |  |
| WO       | ro 2004012776 |     |      |     | A1   |      | 2004 | 0212 | 1   | WO 2 | 003- | EP50 | 340 |     | 20030728 |      |     |  |
|          | W:            | ΑE, | AG,  | AL, | AM,  | AT,  | AU,  | ΑZ,  | BA, | BB,  | BG,  | BR,  | ΒY, | ВŻ, | CA,      | CH,  | CN, |  |
|          |               | co, | CR,  | Cυ, | CZ,  | DE,  | DK,  | DM,  | DZ, | EC,  | EE,  | ES,  | FI, | GB, | GD,      | GΕ,  | GH, |  |
|          |               | GΜ, | HR,  | HU, | ID,  | IL,  | IN,  | IS,  | JP, | ΚE,  | KG,  | KP,  | KR, | ΚZ, | LC,      | LK,  | LR, |  |
|          |               | LS, | LT,  | LU, | LV,  | MA,  | MD,  | MG,  | MK, | MON, | MW,  | MΧ,  | ΜZ, | NI, | NO,      | ΝZ,  | OM, |  |
|          |               | PH, | PL,  | PT, | RO,  | RU,  | sc,  | SD,  | SE, | SG,  | sĸ,  | SL,  | ΤJ, | TM, | TN,      | TR,  | TT, |  |
|          |               | TZ, | UA,  | υG, | US,  | υz,  | vc,  | VN,  | YU, | ZA,  | ZM,  | ZW   |     |     |          |      |     |  |
|          | RW:           | GH, | GΜ,  | KE, | LS,  | MW,  | MZ,  | SD,  | SL, | SZ,  | TZ,  | UG,  | ZM, | ZW, | AM,      | A2,  | BY, |  |
|          |               | KG, | ΚZ,  | MD, | RU,  | TJ,  | TM,  | AT,  | BE, | BG,  | CH,  | CY,  | CZ, | DE, | DK,      | EE,  | ES, |  |
|          |               | FI, | FR,  | GB, | GR,  | Hυ,  | IE,  | IT,  | LU, | MC,  | NL,  | PT,  | RO, | SE, | SI,      | sĸ,  | TR, |  |
|          |               | BF, | ВJ,  | CF, | CG,  | CI,  | CM,  | GΑ,  | GΝ, | GQ,  | G₩,  | ML,  | MR. | NE, | ŚN,      | TD,  | TG  |  |
| PRIORITY | APP           | LN. | info | . : |      |      |      |      |     | EP 2 | 002- | 7816 | 4   |     | A 2      | 0020 | 801 |  |

OTHER SOURCE(S):

MARPAT 140:163705

This invention pertains to a method for producing isotopically labeled [14C] indolinone derivs. With general formula of I [wherein R = alkyl, alkoxy, or halo; Rl = (un)substituted alkyl or coNH2; m = 0.4; n = 0.3] ΑВ OE

pharmaceutically acceptable salts. For example, H14CONMe2 was reacted with 2,4-dimethylpyrrole in diphosphoryl chloride to give 3,5-dimethyl-1H-pyrrole-2-[14C]carboxaldehyde (49%). The above aldehyde was reacted with oxindole in EtOH in the presence of pyrrolidine to afford

II (54%). 656253-79-1P 656253-81-5P 656253-83-7P RL: SPN (Synthetic preparation); PREP (Preparation)

ANSWER 23 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

L4 ANSWER 23 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
(prepn. of isotopically labeled indolinone derivs.)

RN 65623-79-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl-14C]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown

656253-81-5 CAPLUS 1H-Pyrrole-3-carboxamide,  $5-\{(z)-\{5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl-14C]-N-\{(2S)-2-hydroxy-3-(4-morpholinyl)propyl]-2, 4-dimethyl- (9CI) (CA INDEX NAME)$ 

Absolute stereochemistry. Double bond geometry as shown.

656253-83-7 CAPLUS

1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[[5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl-14C]-2,4-dimethyl- [9CI] (CA INDEX NAME)

L4 ANSWER 24 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2004:100803 CAPLUS
DOCUMENT NUMBER: 140:139483
TITLE: Method for enhancing the effectiveness of therapies of

INVENTOR(S):
PATENT ASSIGNEE(S):
SOURCE:

hyperproliferative diseases
Chang, Yan; Sasak, Vodek
USA
US. Pat. Appl. Publ., 14 pp., Cont.-in-part of U.S.
Ser. No. 176, 235.
CODEN: USXXCO
Patent
English
3

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| PATENT NO. |                 |     |    |     |      |      | DATE |      |     |      |        |       |       | DATE  |       |      |     |  |
|------------|-----------------|-----|----|-----|------|------|------|------|-----|------|--------|-------|-------|-------|-------|------|-----|--|
|            | S 2004023925 A1 |     |    |     |      |      | 2004 | 0205 |     |      | 2003-  |       |       |       |       |      |     |  |
|            | 2003            |     |    |     |      |      |      |      |     |      | 2002-  |       |       |       |       | 0020 |     |  |
| us         | 6680            | 306 |    |     | B2   |      |      | 0120 |     |      |        |       |       |       | _     |      |     |  |
| us         | US 2004043962   |     |    |     |      |      |      |      |     | US   | 2003-  | 6573  | 83    |       | 2     | 0030 | 808 |  |
|            | WO 2004091634   |     |    |     |      |      |      |      |     |      |        |       |       |       |       |      |     |  |
|            |                 |     |    |     |      |      |      |      |     |      | , BG,  |       |       |       |       |      |     |  |
|            |                 |     |    |     |      |      |      |      |     |      | , EC,  |       |       |       |       |      |     |  |
|            |                 |     |    |     |      |      |      |      |     |      | , JP,  |       |       |       |       |      |     |  |
|            |                 |     |    |     |      |      |      |      |     |      | , MK,  |       |       |       |       |      |     |  |
|            |                 |     |    |     |      |      |      |      |     |      | , sc,  |       |       |       |       |      |     |  |
|            |                 |     |    |     |      |      |      |      |     |      | , UZ,  |       |       |       |       |      |     |  |
|            | RW:             |     |    |     |      |      |      |      |     |      | , SZ,  |       |       |       |       |      |     |  |
|            |                 |     |    |     |      |      |      |      |     |      | , BG,  |       |       |       |       |      |     |  |
|            |                 |     |    |     |      |      |      |      |     |      | , MC,  |       |       |       |       |      |     |  |
|            |                 |     |    |     |      |      |      |      |     |      | GN,    |       |       |       |       |      |     |  |
|            |                 | TD, |    | ٠., | υ,   | ٠.,  | ٠٠,  | ٠.,  | ٠., | •    | , 011, | Ος,   | un,   | 1127  | ruc,  | 146, | BN, |  |
| PRIORITY   | APP             |     |    | .:  |      |      |      |      | ,   | US : | 2001-  | 2999  | 91P   | 1     | 2 2   | 0010 | 621 |  |
|            |                 |     |    |     |      |      |      |      | ,   | US : | 2002-  | 1762  | 35    | ,     | 12 20 | 0020 | 520 |  |
|            |                 |     |    |     |      |      |      |      | 1   | US : | 2003-  | 1087  | 23    | ,     | A 20  | 0030 | 107 |  |
|            |                 |     |    |     |      |      |      |      | ,   | us : | 2003-  | 4610  | 06P   | 1     | 2 2   | 0030 | 107 |  |
|            |                 |     |    |     |      |      |      |      | ,   | JS : | 2003-4 | 4745  | 62 P  | 1     | 2 20  | 0030 | 530 |  |
| AB The     | eff             | cac | of | con | ent: | iona | l ca | ncer | the | rap: | ies sı | ich a | as s: | ırgei | ry,   |      |     |  |

The efficacy of conventional cancer therapies such as surgery, chemotherapy and radiation is enhanced by the use of a therapeutic material which binds to and interacts with galectins. The therapeutic material can enhance apoptosis thereby increasing the effectiveness of oncolytic agents. It can also inhibit angiogenesis thereby moderating tumor growth and/or metastasis.

537795-19-4, SU11248
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(method for enhancing effectiveness of therapies of hyperproliferative diseases)

diseases)
537795-19-4
HH-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

10081147

L4 ANSWER 24 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN Double bond geometry as shown. (Continued)

ANSWER 25 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

REFERENCE COUNT:

THERE ARE  $\theta$  CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

L4 ANSWER 25 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2004:92033 CAPLUS
DOCUMENT NUMBER: 141:94116
Powder-in-Bottle Formulation of SU011248. Enabling Rapid Progression into Human Clinical Trials
Sistla. Anand: Sunga, Alan; Phung, Kenneth; Koparkar, Arun; Shenoy, Narmada
CORPORATE SOURCE: Pharmacia Company, Sugen Inc., San Francisco, CA, 94080, USA
SOURCE: Drug Development and Industrial Pharmacy (2004), 30(1), 19-25
CODEN: DDIPBG; ISSN: 0363-9045
Marcel Dekker, Inc.
DOCUMENT TYPE: Journal
LANGUAGE: English
AB SU011248 is an oral, multitargeted receptor tyrosine kinase inhibitor (anti PDGFR, VEGFR, Kit, and Flt3) for the treatment of solid tumors.

The powder-in-bottle (PIB) approach was used to accelerate development and introduction into Phase I clin. trials. This approach consists of extemporaneously compounding the active pharmaceutical ingredient (API) into a solution or a suspension in the clinic prior to oral administration.

The development consisted of physico-chemical assessment, constitution fluid selection, weighter and development.

selection, weighing and dosing validation, and stability evaluation of API, before and after constitution with the fluid. Of the oral liqs. evaluated, apple juice was selected as the constitution fluid. Particle size of SU011248 had an impact on the weighing validation and the

size of SUULIZE mas an amportant of the dissoln.

dissoln.

time. Particle size specifications of breadth d90<180 µm and length d90<750 µm were set to achieve pharmaceutical acceptability. Dosing validation studies showed complete recovery of SU011248 from the bottle over a dose range of 10 to 2200 mg. SU011248 is stable as the solid API. Following constitution with apple juice, the product is stable through

predicted duration of compounding and dosing at the clin. site. This approach provided a high degree of dosing flexibility during the initial phase of clin. trials. Addnl., the PIB approach reduced the time and API required for clin. development and supplies to < 2 mo and < 100 gm, resp. 557795-19-4, SU 11248

ΙT

S57795-19-4, SU 11248
RL: PKT (Pharmacokinetics); PRP (Properties); BIOL (Biological study) (powder-in-bottle formulation of SU011248. enabling rapid progression into human clin. trials)
557795-19-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAMPY)

Double bond geometry as shown.

L4 ANSWER 26 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2004:41213 CAPLUS

TITLE: 100:105249

Combination of mTOR inhibitor and a tyrosine kinase inhibitor for the treatment of neoplasms

Neel, Benjamin G.; Mohi, Golam

Beth Israel Deaconess Medical Center, USA PCT Int. Appl., 63 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| PATENT NO. |               |     |      |     |     | KIND DATE   |      |      |     | APPL | DATE |             |     |     |          |      |     |  |
|------------|---------------|-----|------|-----|-----|-------------|------|------|-----|------|------|-------------|-----|-----|----------|------|-----|--|
|            |               |     |      |     |     |             |      |      |     |      |      |             |     |     |          |      |     |  |
| WO         | WO 2004004644 |     |      |     |     | A2 20040115 |      |      |     | WO 2 | 003- | <b>US20</b> | 972 |     | 20030703 |      |     |  |
| WO         | WO 2004004644 |     |      |     | A3  |             | 2004 | 0506 |     |      |      |             |     |     |          |      |     |  |
|            | W:            | ΑE, | AG,  | AL, | AM, | AT,         | AU,  | AZ,  | BA, | BB,  | BG,  | BR,         | BY, | BZ, | CA,      | CH,  | CN, |  |
|            |               | co, | CR,  | Cυ, | CZ, | DE,         | DK,  | DM,  | DZ, | EC,  | EE,  | ES,         | FI, | GB, | GD,      | GE.  | GH, |  |
|            |               | GM, | HR,  | ΗU, | ID, | IL,         | IN,  | IS,  | JP, | KE,  | KG,  | KP,         | KR, | KZ, | LC,      | LK.  | LR. |  |
|            |               | LS, | LT,  | LU, | LV, | MA,         | MD,  | MG,  | MK, | MN,  | MW,  | MX,         | MZ, | NI, | NO.      | NZ.  | OM, |  |
|            |               | PG, | PH,  | PL, | PT, | RO,         | RU,  | SC,  | SD, | SE,  | SG,  | SK,         | SL, | SY, | TJ,      | TM.  | TN. |  |
|            |               | TR, | TT,  | TZ, | UA, | UG,         | US,  | UZ,  | VC, | VN,  | YU,  | ZA,         | ZM, | ZW  |          |      |     |  |
|            | RW:           | GH, | GΜ,  | ΚE, | LS, | MW,         | MZ,  | SD,  | SL, | SZ,  | TZ,  | UG,         | ZM, | ZW, | AM,      | AZ,  | BY, |  |
|            |               | KG, | ΚZ,  | MD, | RU, | TJ,         | TM,  | ΑŤ,  | BE, | BG,  | CH,  | CY,         | CZ, | DE, | DK,      | EE,  | ES, |  |
|            |               | FI, | FR,  | GB, | GR, | ΗU,         | ΙE,  | IT,  | LU, | MC,  | NL,  | PT,         | RO, | SE, | SI,      | SK,  | TR, |  |
|            |               | BF, | ВJ,  | CF, | CG, | CI,         | CM,  | GΑ,  | GN, | GQ,  | GW,  | ML,         | MR, | NE, | SN,      | TD,  | TG  |  |
| PRIORITY   | APP           | LN. | INFO | .:  |     |             |      |      |     | US 2 | 002- | 3940        | 29P |     | P 2      | 0020 | 705 |  |

The invention features methods and compns. including an mTOR inhibitor

a tyrosine kinase inhibitor for reducing the proliferation of and enhancing the apoptosis of neoplastic cells. The addition of an MEK inhibitor to this combination further enhances the effectiveness of this

US 2002-412402P

therapeutic method.
SST/95-19-4, SU11248
SR: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(combination of mTOR inhibitor and tyrosine kinase inhibitor for

therapy)
557795-19-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-{2-(diethylamino)ethyl]-5-{(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown

P 20020920

L4 ANSWER 26 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

(Continued)

ANSWER 27 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

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ANSWER 27 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
SSION NUMBER: 2004:20448 CAPLUS
  L4 ANSWER 27 OF
ACCESSION NUMBER:
  DOCUMENT NUMBER:
                                                                          140:87676
   TITLE:
                                                                          Derivatives of gambogic acid and analogs as
  activators
                                                                         of caspases and inducers of apoptosis
Tseng, Ben; Sirisoma, Nilantha Sudath; Cai, Sui
  INVENTOR (S):
                                                                          Zhang, Han-Zhong; Kasibhatla, Shailaja; Ollis,
  Kristin
                                                                        P.: Drewe, John A.
Cytovia, Inc., USA
PCT Int. Appl., 92 pp.
CODEN: PIXXD2
Patent
English
 PATENT ASSIGNEE(S):
SOURCE:
 DOCUMENT TYPE:
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                PATENT NO.
                                                                         KIND
                                                                                         DATE
                                                                                                                               APPLICATION NO.
PATENT NO. KIND DATE APPLICATION NO. DATE

100 2004002428 A2 20040108 W0 2003-US20668 20030701

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GH, HR, HU, ID, IL, IN, IS, JP, KE, KG, KF, KR, KZ, LC, LK, LR, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, HZ, NI, NO, NZ, OM, PG, PH, PI, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG US 2004082066 A1 20040429 US 2003-608670 20030701

PRIORITY APPLN. INFO:
                                                                                                                               US 2002-413649P
                                                                                                                                                                                         P 20020926
  OTHER SOURCE(S): MARPAT 140:87676

AB The invention is directed to derivs. of gambogic acid and analogs thereof.
               Exemplary gambogic acid derivs. of the present invention include, among others, derivs. substituted in the C10 and C28 positions of gambogic
                The present invention also relates to the discovery that certain
          offerred compds. of the invention are activators of caspases and inducers of apoptosis. Therefore, the activators of caspases and inducers of apoptosis of this invention can be used to induce cell death in a variety of clin. conditions in which uncontrolled growth and spread of abnormal cells occurs.

557795-19-4, SU11248
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(derivs. of gambogic acid and analogs as activators of caspases and inducers of apoptosis)
557795-19-4 CAPLUS
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L4 ANSWER 28 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2003:967557 CAPLUS
DOCUMENT NUMBER: 140:174343
TITLE: Quantitation of SU11248, an oral multi-target
tyrosine
                                                                         kinase inhibitor, and its metabolite in monkey
                                                                       by liquid chromatograph with tandem mass spectrometry following semi-automated liquid-liquid extraction Baratte, S.; Sarati, S.; Frigerio, E.; James, C. A.; Ye, C.; Zhang, Q. Global Drug Metabolism, Nerviano, 20014, Italy Journal of Chromatography, A (2004), 1024(1-2), 87-94 CODEN: SCRAEY; ISSN. 0021-9673 Elsevier Science B.V. Journal
  AUTHOR (S):
   CORPORATE SOURCE:
SOURCE:
PUBLISHER: Elsevier Science B.V.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB SUll248 is a potent inhibitor of PDGFR, VEGFR, KIT, and Flt3, and is currently under Phase I clin. evaluation as an anticancer drug. A sensitive and specific anal. method for the quantitation of SUll248 and its metabolite in several monkey tissues (liver, kidney, brain and white fat) using LC-MS-MS following semi-automated liquid-liquid extraction (LLE) was
                was developed and validated. Amts. of 50 mg of tissue were homogenized using an ultrasonic processor. After addition of the stable labeled internal
  standard
(IS) and ammonium hydroxide (0.3%), samples were extracted with 2.5 mL of
tert-Bu Me ether. Following centrifugation, aliquots of 1.8 mL of the
organic phase were transferred into a 96-well plate. The Packard
                II robotic liquid handler was used to perform all steps mentioned above.
The organic phase was dried and the residue was reconstituted with 800
                of 15 mM ammonium formate buffer solution (pH 3.25) using a Tomtec
  Quadra 96
                workstation. Aliquots of 10 µL of the resulting solution were injected into the LC-MS-MS system. A Symmetry Shield C8 column (50 mm+2.1 mm, 3.5 µm) was used to perform the chromatog, anal. The mobile phase was 15 mM ammonium formate buffer solution (pH 3.25)-MeCN (74:26
  pos. ion mode. The method was validated for both compds. over the calibration range of .apprx.2 and 2000 ng/g. The suitability and robustness of the method for in vivo samples were confirmed by anal. of monkey tissues from animals dosed with SUI1248.

557795-19-4. SUI1248 557795-19-4D, SUI1248, metabolite
RL: ANT (Analyte); ANST (Analytical study)
(quantitation of SUI1248, an oral multi-terget tyrosine kinase inhibitor, and its metabolite in monkey tissues by liquid matocraph
  Chromatograph
with tandem mass spectrometry following semi-automated liquid-liquid
 extraction tendem mass spectrometry following semi-automated liquid-liquid
RN 557795-19-4 CAPLUS

IH-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-
dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX
NAME)
                  Page 28
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10081147

L4 ANSWER 28 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

557795-19-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-{2-(diethylemino)ethyl}-5-{(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

REFERENCE COUNT: THIS

THERE ARE 10 CITED REFERENCES AVAILABLE FOR 10

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L4 ANSWER 29 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN Double bond geometry as shown.

NEt2

REFERENCE COUNT:

THERE ARE 38 CITED REFERENCES AVAILABLE FOR

FORMAT

RECORD. ALL CITATIONS AVAILABLE IN THE RE

L4 ANSWER 29 OF 65
ACCESSION NUMBER:
2003:949777 CAPLUS
DOCUMENT NUMBER:
141:64493
SUI1248 inhibits tumor growth and CSF-1R-dependent osteolysis in an experimental breast cancer bone metastasis model
AUTHOR(S):

AUTHOR(S):

AUTHOR(S):

AUTHOR(S):

CORPORATE SOURCE:

CORPORATE SO

PUBLISHER: Kluwer Academic Publishers
DOCUMENT TYPE: Journal
LANGUAGE: Brglish
B The aim of the study was to investigate inhibitory effects of the
receptor

tyrosine kinase (RTK) inhibitor SU11248 against CSF-1R and osteoclast

formation. We developed an in vivo model of breast cancer metastasis to evaluate efficacy of SU11248 against tumor growth and tumor-induced osteolysis in bone. The in vitro effects of SU11248 on CSF-IR phosphorylation, OC formation and function were evaluated. Effects on 435/HAL-Luc tumor growth in bone were monitored by in vivo

mminescence imaging (BLI), and inhibition of osteolysis was evaluated by measurement of serum pyridinoline (PYD) concentration and histol. Phosphorylation

ne receptor for M-CSF (CSF-1R) expressed by NIH3T3 cells was inhibited by SU11248 with an ICSO of 50-100 nM, consistent with CSF-1R belonging to

class III split kinase domain RTK family. The early M-CSF-dependent

phase of in vitro murine OC development and function were inhibited by SU11248 at 10-100 nM. In vivo inhibition of osteolysis was confirmed by significant lowering of serum PYD levels following SU11248 treatment of tumor-bearing mice (P=0.047). Using BLI, SU11248 treatment at 40 mg/kg/day for 21 days showed 64% inhibition of tumor growth in bone (P=0.069), and at 80 mg/kg/day showed 89% inhibition (P=0.001). Collectively, these data suggest that SU1124% may be an effective and tolerated therapy to inhibit growth of breast cancer bone metastases,

with

the addnl. advantage of inhibiting tumor-associated osteolysis. 557795-19-4, SU 11248 RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (SU11248 inhibits tumor growth and CSF-1R-dependent osteolysis in

mouse

breast cancer bone metastasis model and mechanisms involved)
557795-19-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-({Z}-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

L4 ANSWER 30 OF 65
ACCESSION NUMBER:
DOCUMENT NUMBER:
11:33400
Proof of Target for SUl1654: Inhibition of KIT
Phosphorylation in Canine Mast Cell Tumors
Pryer, Nancy K.; Lee, Lesile B.; Zadovaskaya, Regina;
Yu, Xiaoming; Sukbuntherng, Juthamas; Cherrington,
Julie M.; London, Cheryl A.
SUGEN, Inc, South San Francisco, CA, USA
COEDE:
COEDE: CREF4; ISSN: 1078-0432
American Association for Cancer Research
JOULMENT TYPE:
JOURNALL SUGENCE CALLED SUGENCE CALL

(SHER: American Association for Lancer Research
MENT TYPE: Journal
JAGE: English
PURPOSS: The purpose of this study was to evaluate the effect of the
receptor tyrosine kinase inhibitor SU11654 on the activity of its mol.
target KIT in canine mast cell tumors (MCT) and correlate target
inhibition with mutational status of the c-kit juxtamembrane domain and
SU11654 plasma concentration Exptl. Design: Tumor biopsies were

obtained from

ined from dogs with advanced MCTs before and 8 h after administration of a single oral dose of SU11654, previously shown to be active in dogs with MCTs. Blood samples were taken to determine the plasma concentration of 554. Levels of

Blood samples were taken to determine the plasma concentration of 654. Levels of phosphorylated KIT and ERKI/2 were assessed in tumor biopsies by Western blot. Tumors were analyzed by PCR for the presence or absence of an internal tandem duplication (ITD) in the juxtamembrane domain of c-kit. RESULTS: Fourteen dogs with advanced MCTs were enrolled in the study; 10 of these were evaluable for KIT target modulation (the remaining tumor specimens had inevaluable amts. of total KIT protein). Of these, eight MCTs showed reduced levels of phosphorylated KIT relative to total KIT after treatment with SU11654, compared with pretreatment biopsies. All four evaluable MCTs expressing ITD mutant c-kit showed modulation of KIT phosphorylation, as did four of seven tumors expressing non-ITD c-kit. Phosphorylated ERKI/2 was modulated in seven tumors; this did not correlate with inhibition of KIT phosphorylation. CONCLUSION: SU11654 treatment at the efficacious dose results in inhibition of KIT phosphorylation in canine MCTs.
350068-94-5, SU11654

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES

(SU11654 effect on activity of mol. target KIT in canine mast cell

tumors)

CAPLUS

1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-(2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX

Double bond geometry as shown.

Page 29 SAEED L4 ANSWER 30 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

REFERENCE COUNT:

44 THERE ARE 44 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

ANSWER 31 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) anti-FLT3 activity in patients. Proof of target inhibition accomplishes

IT

crucial milestone in the development of novel oncol. therapeutics. 557795-19-4, SU 11248
RL: ADV (Adverse effect, including toxicity); DMA (Drug mechanism of action); PAC (Pharmacological activity); BIOL (Biological study)
(SU 11248 FLT3 phosphorylation inhibition in acute myeloid leukemia patients) 557795-19-4

patients)
55795-19-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

Double bond geometry as shown.

REFERENCE COUNT:

34 THERE ARE 34 CITED REFERENCES AVAILABLE FOR

FORMAT

RECORD. ALL CITATIONS AVAILABLE IN THE RE

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L4 ANSWER 31 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2003:943061 CAPLUS
DOCUMENT NUMBER: 140:399449
An Innovative Phase I Clinical Study Demonstrative Phase I Clinical Study Demon
```

Myeloid Leukemia Patients O'Farrell, Anne-Marie; Foran, James M.; Fiedler, Walter; Serve, Hubert: Paquette, Ron L.; Cooper, Maureen A.; Yuen, Helene A.; Louie, Sharianne G.; AUTHOR (S): Kim,

Heidi; Nicholas, Susan; Heinrich, Michael C.; Berdel, Wolfgang E.; Bello, Carlo; Jacobs, Mark; Scigalla, Paul; Manning, William C.; Kelsey, Stephen; Cherrington, Julie M. SUGEN Inc., South San Francisco, CA, USA Clinical Cancer Research (2003), 9(15), 5465-5476 CODEN: CCRF4; ISSN: 1078-0422 American Association for Cancer Research Journal

CORPORATE SOURCE:

PUBLISHER:

Journal English

DOCUMENT TYPE: LANGUAGE: AB PURPOSE: 0 DAGE: Engilen
PURPOSE: Obtaining direct and rapid proof of mol. activity in early clin.
trials is critical for optimal clin. development of novel targeted

trials is critical for optimal Call. General Management with selectivity for SU 11248 is an oral multitargeted kinase inhibitor with selectivity for fms-related tyrosine kinase 3/F1k2 (FLT3), platelet-derived growth factor receptor a/B, vascular endothelial growth factor receptor 1/2, and KIT receptor tyrosine kinases. FLT3 is a promising candidate for targeted therapy in acute myeloid leukemia (AML), because activating mutations occur in up to 30% of patients. We conducted an innovative single-dose clin. study with a primary objective to demonstrate inhibition

of FLT3 phosphorylation by SU 11248 in AML. Exptl. Design: Twenty-nine AML patients each received a single dose of SU 11248, escalated from 50 to

350 mg, in increments of 50 mg and cohorts of three to six patients. FLT3

phosphorylation and plasma pharmacokinetics were evaluated at seven time points over 48 h after SN 11248 administration, and FLT3 genotype was determined Study drug-related adverse events occurred in 31% of

determined Study drug-related education.

determined Study drug-related education.

mainly grade 1 or 2 diarrhea and nausea, at higher dose levels. RESULTS: Inhibition of FLT3 phosphorylation was apparent in 50% of FLT3-wild-type (WT) patients and in 100% of FLT3-mutant patients. FLT3 internal tandem duplication (ITD) mutants showed increased sensitivity relative to FLT3-WT, consistent with preclin. predictions. The primary end point, strong inhibition of FLT3 phosphorylation in >50% patients, was reached in RESULTS:

200 mg and higher dose cohorts. Downstream signaling pathways were also inhibited; signal transducer and activator of transcription 5 (STAT5) was reduced primarily in internal tandem duplication patients and at late

points in FLT3-WT patients, whereas extracellular signal-regulated kinase (ERK) activity was reduced in the majority of patients, independent of FLT3 inhibition. CONCLUSIONS: This novel translational study bridges preclin. models to the patient setting and provides the first evidence of

```
L4 ANSWER 32 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN ACCESSION NUMBER: 2003:931518 CAPLUS DOCUMENT NUMBER: 140:689
```

DOCUMENT NUMBER: TITLE:

140:689
Genes showing altered patterns of expression in response to inhibition of tyrosine kinases and their use in screening kinase inhibitors
Morimoto, Alyssa; Deprimo, Samuel; O'Farrell,
Anne-Marie; Smolich, Beverly Dp.; Manning, William C.;
Walter, Sarah A.; Schilling, James Walter, Jr.;
Cherrington, Julie
Sugen, Inc., USA
PCT Int. Appl., 408 pp.
CODEN: PIXXD2

INVENTOR (S):

PATENT ASSIGNEE(S): SOURCE:

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| PA            | KIND DATE |      |     |     |             | APPL | ICAT | DATE            |     |      |      |      |          |     |          |      |     |  |
|---------------|-----------|------|-----|-----|-------------|------|------|-----------------|-----|------|------|------|----------|-----|----------|------|-----|--|
|               |           |      |     |     |             |      |      |                 |     |      |      |      |          |     |          |      |     |  |
| WO 2003097854 |           |      |     |     | A2 20031127 |      |      | WO 2003-US15711 |     |      |      |      |          |     | 20030519 |      |     |  |
|               | W:        | ΑE,  | AG, | AL, | AM,         | AT,  | ΑU,  | ΑZ,             | BA, | BB,  | BG,  | BR,  | BY,      | BZ, | CA,      | CH,  | CN, |  |
|               |           | co,  | CR, | cu, | CZ,         | DE,  | DK,  | DM,             | DZ, | EC,  | EE,  | ES,  | FI,      | GB, | GD,      | GE,  | GH, |  |
|               |           | GΜ,  | HR, | HU, | ID,         | IL,  | IN,  | IS,             | JP, | KE,  | KG,  | KP,  | KR,      | KZ, | LC,      | LK,  | LR, |  |
|               |           | LS,  | LT, | LU, | LV,         | MA,  | MD,  | MG,             | MK, | MN,  | MW,  | MX,  | MZ,      | NI, | NO,      | NZ,  | OM, |  |
|               |           | PH,  | PL, | PT, | RO,         | RU,  | SC,  | SD,             | SE, | SG,  | SK,  | SL,  | TJ,      | TM, | TN.      | TR.  | TT. |  |
|               |           | TZ,  | UA, | UG, | US,         | UZ,  | VC,  | VN,             | YU, | ZA,  | ZM,  | ZW   |          |     |          |      |     |  |
|               | RW:       | GH,  | GM, | KE, | LS,         | MW,  | MZ,  | SD,             | SL, | SZ,  | TZ,  | UG,  | ZM,      | ZW, | AM,      | AZ,  | BY, |  |
|               |           | KG,  | KZ, | MD, | RU,         | TJ,  | TM,  | AT,             | BE, | BG,  | CH,  | CY,  | CZ,      | DE, | DK,      | EE,  | ES. |  |
|               |           | FI,  | FR, | GB, | GR,         | HU,  | IE,  | IT,             | LU, | MC,  | NL,  | PT,  | RO,      | SE, | SI,      | SK,  | TR. |  |
|               |           | BF,  | ВJ, | CF, | CG,         | CI,  | CM,  | GA,             | GN, | GQ,  | GW,  | ML,  | MR.      | NE, | SN,      | TD.  | TG  |  |
| US            | 2004      | 0185 | 28  |     | A1          |      | 2004 | 0129            |     | US 2 | 003- |      | 20030519 |     |          |      |     |  |
| PRIORIT       |           |      |     |     |             | US 2 | 002- | 3808.           | 72P |      | P 2  | 0020 | 517      |     |          |      |     |  |
|               |           |      |     |     |             |      |      |                 |     | US 2 | 003- | 4488 | 74P      |     | P 2      | 0030 | 224 |  |

OTHER SOURCE(S):

R SOURCE(S): MARPAT 140:689
Genes that are regulated by tyrosine kinase-dependent signal transduction pathways are identified as markers for the screening of inhibitors of kinase activity. The change in levels of either the protein or mRNA in a suitable test system may be used to assess the effectiveness of a test compound as an inhibitor of a tyrosine kinase activity. The invention

US 2003-448922P

relates to novel methods, wherein a change in the level of at least one biomarker in a mammal exposed to a compound, compared to the level of the biomarker(s) in a mammal that has not been exposed to the compound, indicates whether the mammal is being exposed to, or is experiencing or will experience a therapeutic or toxic effect in response to, a compound that inhibit tyrosine kinase activity.

342641-63-8 342641-64-9 342641-94-5
346405-32-1 35608-92-1 35608-90-1
515138-82-6 627908-83-2 627908-94-6
627908-92-3 627908-93-4 627908-97-6
627908-92-3 627908-93-4 627908-94-5
627908-92-6

BL: BSU (Biological study, unclassified); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (as tyrosine kinase inhibitor; genes showing altered patterns of

P 20030224

## 10081147

ANSWER 32 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) expression in response to inhibition of tyrosine kinases and their use in screening kinase inhibitors) 342641-63-8 CAPLUS (APPLICATION OF THE PRINCIPLE OF THE PRINCIP

342641-64-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA

342641-94-5 CAPLUS 1H-Pyrrole-3-carboxamide, N-(2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

ANSWER 32 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356068-90-1 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

515138-82-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 32 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

346405-32-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA

RN 356068-82-1 CAPLUS
CN 1H-Fyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX..... NAME)

ANSWER 32 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

RN 627908-83-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-[(1,2-dihydro-5-iodo2-0xo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

627908-84-3 CAPLUS

IH-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-5-iodo-2-oxo-3H-indol-3ylidene)methyl-2,4-dimethyl-N-{2-(1-pyrrolidinyl)ethyl}- (9CI) (CA INDEX NAME)

L4 ANSWER 32 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 627908-85-4 CAPLUS
IH-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3ylidene)methyl)-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAME)

RN 627908-86-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-{2-(4-morpholinyl)ethyl}- (9CI) (CA INDEX NAME)

L4 ANSWER 32 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued Double bond geometry unknown.

RN 627908-93-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-N-[(2S)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

RN 627908-94-5 CAPLUS
CN | H-Pytrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-yliden)methyl]-N-[(25)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown

RN 627908-95-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-5-iodo-2-oxo-3H-indol-3-ylidene)methyl)---[(25)-2-hydroxy-3-(4-morpholinyl)propyl)-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

L4 ANSWER 32 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 627908-87-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-5-iodo-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAMZ)

RN 627908-92-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3ylidene)methyl]-N-[(25)-2-hydroxy-3-{4-morpholinyl)propyl}-2,4-dimethyl(9C1) (CA INDEX NAME)

Absolute stereochemistry.

L4 ANSWER 32 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued

Page 32 SAEED

10081147

L4 ANSWER 33 OF 65
ACCESSION NUMBER:
DOCUMENT NUMBER:
140:122316
Hair depigmentation is a biological readout for pharmacological inhibition of KIT in mice and humans Moss, Katherine G.; Toner, Guy C.; Cherrington, Julie M.; Mendel, Dirk B.; Laird, A. Douglas
Department of Preclinical Research and Exploratory Development, SUGEN, Inc., South San Francisco, CA,

USA SOURCE:

Development, SUGEN, Inc., South San Francisco, CA,

RCE: Journal of Pharmacology and Experimental Therapeutics
(2003), 307(2), 476-488
CODEN: JPETAB; ISSN. 0022-3565
American Society for Pharmacology and Experimental
Therapeutics
JOURNT TYPE: Journal
English
Deregulated activation of the KIT receptor tyrosine kinase has been
implicated in several human cancers and in inflammation, making it an
attractive target for therapeutic intervention. Conversely, deficiencies
in KIT signaling have been implicated in human and-animal hair
pigmentation disorders, reflecting a role for KIT in the development and
function of melanocytes. The goal of this study was to explore the
potential utility of hair depigmentation as a biol. readout for systemic
inhibition of KIT by SU11248 5-[5-fluor-2-oxo-1,2-dihydroindol-(32)ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
(2-diethylaminoethylamide), an oral multitargeted tyrosine kinase
inhibitor with antitumor and antiangiogenic activity through targeting
platelet-derived growth factor receptors, vascular endothelial growth
factor receptors, KIT, and FLT3. Oral SU11248 treatment induced
dose-dependent depigmentation of newly regrown hair in depliated C57BL/6
mice. Similar effects were seen after administration of a
KIT-neutralizing antibody. SU11248-induced hair depigmentation was
reversible with cessation of treatment. Histol. and immunohistochem.
evaluation of mouse skin samples supported these observations and
saled

that SU11248 has no effect on levels of KIT-pos. melanocytes associated

hair follicles, indicating that the inhibitory effect is at the level of melanocyte function rather than their development/survival. Similar hair depigmentation has been noted in several cancer patients receiving

in phase I trials. Strikingly, patient scalp hair exhibits bands of depigmentation and pigmentation that correspond, resp., to periods of treatment and dosing rest periods. These data demonstrate that hair pigmentation can serve as a dose-dependent, dynamic, biol. readout for

inhibition in mice, and, apparently, in humans.

557795-19-4, SU11248

RI: ADV (Adverse effect, including toxicity); DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(hair depigmentation is a biol. readout for pharmacol. inhibition of KIT in mice and humans)

557795-19-4 CAPLUS

IH-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(2)-(5-fluoro-1,2-

L4 ANSWER 34 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2003:844931 CAPLUS
DOCUMENT NUMBER: 140:192417
TITLE: Preclinical evaluation of the tyrosine kinase inhibitor SU11248 as a single agent and in

with "standard of care" therapeutic agents for the treatment of breast cancer
Abrams, Tinya J.; Murray, Lesley J.; Pesenti, Enrico; Walker Holway, Vicky; Colombo, Tina; Lee, Leslie B.; Cherrington, Julie M.; Pryer, Nancy K.
Preclinical Research and Experimental Development, SUGEN, Inc., South San Francisco, CA, USA
Molecular Cancer Therapeutics (2003), 2(10), AUTHOR (S):

CORPORATE SOURCE:

CODEN: MCTOCF; ISSN: 1535-7163 American Association for Cancer Research

PUBLISHER: American Association for Cancer Research
DOCUMENT TYPE: Journal
LANGUAGE: English
AB SUI1248 is an oral multitargeted tyrosine kinase inhibitor with antitumor
and antiangiogenic activities through targeting platelet-derived growth
factor receptor, vascular endothelial growth factor receptor, KIT, and
FLT3, the first three of which are expressed in human breast cancer

and/or

its supporting tissues. The purpose of the present studies was to
demonstrate the potent anticancer activity of SU11248 alone or in
combination with conventional cytotoxic agents against several distinct
preclin. models of breast cancer. SU11248 was administered as a
monotherapy to (1) mouse mammary tumor virus-v-Ha-ras mice and
7,12-dimethylbenz(a)anthracene-treated rats bearing mammary tumors and

mice bearing human breast cancer xenografts of s.c. MX-1 tumors and osseous metastasis of a MDA-MB-435-derived cell line (435/HAL-Luc). SU11248 was also administered in combination with docetaxel both in xenograft models and in combination with 5-fluorouracil and doxorubicin

the MX-1 model. SU11248 treatment potently regressed growth of mammary cancers in mouse mammary tumor virus-v-Ha-ras transgenic mice (82% regression) and 7,12-dimethylbenz(a)anthracene-induced mammary tumors in rats (99% regression at the highest dose; P< 0.05 for both). This agent also inhibited MX-1 tumor growth by 52%, with markedly enhanced

cancer effects when administered in combination with docetaxel, 5-fluorouracil, or doxorubicin compared with either agent alone (P < 0.05). SU11248 treatment in combination with docetaxel effectively prolonged survival of mice, with 435/HAL-Luc cancer xenografts established in bone compared

either agent alone (P < 0.05). These results demonstrate that SU11248 is effective in preclin. breast cancer models and suggest that it may be useful in the treatment of breast cancer in the clinic.

557795-19-4, SU11248
RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (antitumor tyrosine kinase inhibitor SU11248 for treatment of breast

cancer)
557795-19-4 CAPLUS
HR-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

ANSWER 33 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

REFERENCE COUNT:

31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L4 ANSWER 34 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN Double bond geometry as shown. (Continued)

REFERENCE COUNT:

30 THERE ARE 30 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

Page 33 SAEED

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L4 ANSWER 35 OF 65
ACCESSION NUMBER:
DOCUMENT NUMBER:
TITLE:
                                                                                                                   CAPLUS COPYRIGHT 2005 ACS on STN
2003:719299 CAPLUS
139:240339
Antitumor agent comprising combination of
sulfonamide-containing heterocyclic compound with
angiogenesis inhibitor
Wakabayashi, Toshiaki; Ono, Naoto; Semba, Taro;
Haneda, Toru
Eisai Co., Ltd., Japan
PCT Int. Appl., 49 pp.
CODEN: PIXXD2
Patent
Japanese
IT: 1
     PATENT ASSIGNEE (S):
SOURCE:
      DOCUMENT TYPE:
     FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                                 PATENT NO.
                                                                                                                                            KIND
                                                                                                                                                                               DATE
                                                                                                                                                                                                                                                 APPLICATION NO.
                                                                                                                                                                                                                                                                                                                                                                             DATE
MO 2003074045

M: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CG, CR, CU, CZ, DE, DK, DE, E, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, VU, ZA, ZM, ZW

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, UT, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TS, TS, KT, TR, BF, BJ, CF, CG, CIT, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

EP 1481678

R: AT, BE, CH, DE, DE, KE, SF, RG, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK

PRIORITY APPLN. INFO::

DATE

200303014

200303034

20030304
```

OTHER SOURCE(S): MARPAT 139:240339

It is intended to provide compns. and kits for treating tumor whereby the angiogenesis inhibitory activity and the antitumor activity of a sulfonamide-containing heterocyclic compound represented by the following

WO 2003-JP2492

W 20030304

L4 ANSWER 36 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2003:678808 CAPLUS
DOCUMENT NUMBER: 139:214331
TITLE: Process for preparing TITLE: Process for preparing
aminocarbonylpyrrolylmethylidene
indolinones from indolinones,
imidazolcarbonylpyrrolecarboxaldehydes, and amines.

Jin, Qingwu; Mauragis, Michael A.; May, Paul D.
PATENT ASSIGNEE(S): Pharmacla & Upjohn Company, USA
COEN: PIXXD2

DOCUMENT TYPE: COEN: PIXXD2
PATENT ASC. NUM. COUNT: Papilish
FAMILY ACC. NUM. COUNT: 2 LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. APPLICATION NO. KIND DATE DATE

PATENT NO. KIND DATE APPLICATION NO.

WO 2003070725 A2 20030828 WO 2003-US4520

WO 2003070725 A3 20040115

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY,
CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI,
GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR,
LS, LT, LU, LY, MA, MD, MG, MK, MN, MM, MX, MZ,
PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM,
UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: GH, GM, KE, LS, MN, MZ, SD, SL, SZ, TZ, UG, ZM,
KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ,
FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE,
BJ, CF, CG, CI, CM, GA, GM, GQ, GW, ML, MR, MC,
CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI,
GM, HR, HU, ID, LI, IN, IS, JP, KE, KG, KP, KR,
LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ,
PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,
UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY,
TJ, TM
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM,
CC, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC,
BF, BJ, CF, CG, CI, CM, GA, GM, GQ, GW, ML, MR,
ER, AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, II, LU,
PRIORITY APPIN. INFO:: 20030214 CA, CH, GD, GE, LC, LK, NZ, OM, TR, TT, BZ, GB, KZ, NO, TN, ZW, AM, AZ, BY, DE, DK, EE, ES, SI, SK, TR, BF, SN, TD, TG 20020215 CA, GD, LC, NZ, TR, KZ, ΒZ, GB, KZ, NO, TN, KG, ZW, NL, NE, AT, BE, CH, PT, SE, TR, SN, TD, TG 20030214 SE, MC, PT, HU, SK US 2002-411732P P 20020918 US 2001-268683P P 20010215 US 2001-312361P WO 2003-US4520

CASREACT 139:214331; MARPAT 139:214331 OTHER SOURCE(S):

ANSWER 35 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) formula (I) can be more effectively exerted. By combining with a VEGF inhibitor or an FGF inhibitor, the sulfonamide-contg. heterocyclic compd. can be effectively employed in treating cancer. S57795-19-4, SUIL1248
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(antitumor agent comprising combination of sulfonamide-containing heterocyclic compound with angiogenesis inhibitora)
557795-19-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-((Z)-(5-fluoro-1,2-dibydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

REFERENCE COUNT: THIS

29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

(R<sup>5</sup>)<sub>P</sub> M#L COR20 AB Title compds. [I; R1-R5 = H, SINYI, NATURE OF THE CONTROL OF TH Title compds. [I; R1-R5 = H, alkyl, alkoxy, cycloalkyl, aryl, be substituted by OH, halo; R8 = H, alkyl; R9 = NR1OR11, OH, COR12, aryl, heterocyclyl containing 1-3 N, S, O, N+(O-)R10, NHCOR13; R10, R11 = H, cyanoalkyl, cycloalkyl, aryl, heterocyclyl containing 1-3 N, S, O; R10R11N (R'-substituted) 5-6 membered heterocyclyl optionally containing 1-3 addnl. N, t. N, O, S; R12 = H, OH, alkoxy, aryloxy; R13 = alkyl, haloalkyl, aralkyl; R', R" = H, alkyl, cyanoalkyl, cycloalkyl, aryl, heterocyclyl containing 1-3 , O; R'R"N = 5-6 membered heterocyclyl optionally containing 1-3 addnl. N, O, S; halo = F, Cl, Br, iodo; J = O, S, NH; l of K, L, M = CCOR6, the others of K, L, M = CR5, CR52, N, NR5, O, S; n, p = 0-2; m = 1-4], were prepared The process comprises reaction of azoles (II) with X2R (R5, J, K, L, M, p are as defined above; Q = CHO, CHS, dioxolanyl, tetrahydrooxazolyl, etc.; X1 = Cl, Br; X2 = H; R = pyrrolyl, thiazolidinethionyl, oxazolidinethionyl, imidazolidinethionyl, proportionally according to the corresponding indolinone. Thus, R2O = OR, R], and reaction of III with IR6 (R6 as defined above) and the corresponding indolinone. Thus, 4-(IH-imidazol-1-ylcarbonyl)-3,5-dimethyl-IH-pyrrole-2-carboxaldehyde, N,N-diethylethelediamine, 5-fluoroxindole, N, O, 1H-pyrrole-2-carboxaidenyue, R.N. Maccon, 18 h at 60° to give 858 N-[2-(diethylamino)ethyl]-5-((2)-(5-fluoro-2-oxo-1,2-dihydro-3H-indol-3-ylidene)methyl)-2,4-dimethyl-1H-pyrrole-3-carboxamide.

IT 326914-12-9P 356068-94-5P 452105-23-6P 452105-24-PP 357795-19-4P 857879-12-7P RI.- IMF (Industrial manufacture); SPN (Synthetic preparation); PREP RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

ANSWER 36 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN R5)p

(process for preparing aminocarbonylpyrrolylmethylideneindolinones from Page 34 SAEED

ANSWER 36 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) indolinones, imidazolcarbonylpyrrolecarboxaldehydes, and amines) 326914-12-9 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(2)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 356068-94-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-((2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX

NAME)

Double bond geometry as shown.

452105-23-6 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-((2S)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(SCI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

ANSWER 36 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

557795-19-4 CAPLUS : 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

587879-12-7 CAPLUS

1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 36 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452105-24-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(2)-{5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-{(2R)-2-hydroxy-3-(4-morpholinyl)propyl)-2,4-dimethyl-9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

452105-25-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(Z)-{5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl-N-{(ZR)-2-hydroxy-3-(4-morpholinyl)propyl)-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

452105-26-9 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-{(ZS)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(SCI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

L4 ANSWER 37 OF 65 CAPLUS COPYRIGHT 2005 ACS ON STN ACCESSION NUMBER: 2003:678506 CAPLUS DOCUMENT NUMBER: 139:214337 TITLE: Preparation 1

Preparation of 3-heteroarylmethylene-1,3-dihydro-2H-

Preparation of 3-neteroaryimethylene-1,3-indol-2-ones as protein kinase inhibitors Lin, Nan-Horng; Sham, Hing L.; Xia, Ping Abbott Laboratories, USA U.S. Pat. Appl. Publ., 10 pp. CODEN: USXXCO Patent English 1

INVENTOR(S):

PATENT ASSIGNEE(S): SOURCE:

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE US 2003162785 US 6797825 PRIORITY APPLN. INFO.: US 2002-317336 A1 B2 20030828 20040928 20021212 US 2001-341410P P 20011213

OTHER SOURCE(S): MARPAT 139:214337

The title compds. [I; X = N, CR3; Y = N, CR4; Z = N, CR5; with the

The compose (I, N = N, cos, I = N, cos, I

10 μM. 550373-92-79 550373-93-89

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(preparation of 3-heteroarylmethylene-1,3-dihydro-2H-indol-2-ones as protein

iein
 kinese inhibitors)
550373-92-7 CAPUUS
2H-Indol-2-one, 3-[[4-[(dimethylamino)methyl]-5-methyl-3-(1-methylethyl)1H-pyrrol-2-yl]methylene]-1,3-dihydro-6-(4-hydroxy-3-methoxyphenyl)-,
(32)- (SCI) (CA INDEX NAME)

Page 35

L4 ANSWER 37 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN Double bond geometry as shown. (Continued)

550373-93-8 CAPLUS
2H-Indol-2-one, 3-[[4-[(dimethylamino)methyl]-5-methyl-3-(1-methylethyl)H-pyrcol-2-yl]methylene]-1,3-dihydro-6-(4-hydroxy-3-methoxyphenyl)-,
(32)-, bis(trifluoroacetate) (salt) (9CI) (CA INDEX NAME)

CRN 550373-92-7 CMF C27 H31 N3 O3

Double bond geometry as shown.

CM 2

CRN 76-05-1 CMF C2 H F3 O2

REFERENCE COUNT: THIS

THERE ARE 41 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L4 ANSWER 38 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2003:652131 CAPLUS
DOCUMENT NUMBER: 139:214237
TITLE: Preparation of nitrate prodrugs able to release

oxide in a controlled and selective way and their use for prevention and treatment of inflammatory,

ischemic and proliferative diseases Scaramuzzino, Giovanni Italy Eur. Pat. Appl., 313 pp. CODEN: EPXXDW Patent English 1

INVENTOR(S):
PATENT ASSIGNEE(S):
SOURCE:

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO. DATE APPLICATION NO. EP 1336602 Al 20030820 EP 2002-425075 20020213
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
PRIORITY APPLN. INFO: EP 2002-425075 20020213

GI

New pharmaceutical compds. of general formula F-(X)q (I) [q=1-5,preferably~l;~F is chosen among drugs such as  $\delta$ -tocopherol, clidanac, diethylhomospermine, glucosamine, thymocartin, vofopitant,

X is chosen among 4 groups M, T, V, and Y where M = ONO2, nitrate salt, nitrite ester, ONO, thoinitrite, SNO, etc., T = OR1-M, ORIORI-M, SRINR2R1-M, NR2R1-M, NR2RISRI-M, etc., R1 = saturated or unsatd., linear

branched alkylene, having 1 to 21 carbon atoms or a saturated or unsatd., optionally heterosubstituted or branched cycloalkylene, having 3 to 7 carbon atoms or an optionally heterosubstituted arylene having 3 to 7 carbon atoms; R2 = H, saturated or unsatd., linear or branched 1-21

carbon atom alkyl, saturated or unsatd. optionally heterosubstituted or branched 3-7

ched 3-7
carbon cycloalkyl, optionally heterosubstituted 3-7 carbon aryl; R1, R2 =
OH, SH, F, Cl, Br, OPO3H2, CO2H, etc.; bond between F and T = carboxylic
ester, carboxylic amide, glycoside, azo, thioester, sulfonic ester, etc.;
V = Z-M2, OZ-M2, NR2Z-M2, R1Z-M2, OR1-M2, OR1Z-M2, M2 = M, R1-M, OR1-M.

L4 ANSWER 37 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

ANSWER 38 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) SR1-M, NR2R1-M; ZM2 = COCH2CH(M2)CH2N+Me3, COCH2CH2COM2, COCH(NRR2)CH2M2, etc.: Y = 4-CCC6H4CH2OMC2, O(CH2)4OND2, COCH(NR2)CH2CMC2, 3-OC6H4CH2OMO2, etc.] were prepd. For example, a-tocopherol reacted with 4-H02CC6H4CH2OMO2 to give the nitroxymethyl deriv. II. The compds. of general formula I are nitrate prodrugs which can release nitric oxide in vivo in a controlled and selective way and without hypotensive side effects and for this reason they are useful for the prepn. of medicines for prevention and treatment of inflammatory, ischemic, degenerative and proliferative diseases of musculoskeletal, tegumental, respiratory, gastrointestinal, genito-urinary and central nervous systems.

S8530-09-6P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of nitrate prodrugs for treating or preventing

(Uses)
(preparation of nitrate prodrugs for treating or preventing inflammatory,
ischemic, degenerative, and proliferative diseases)

RN 586350-09-6 CAPLUS
CN L-Serine, N-[3-[5-[(2)-(1,2-dihydro-2-oxo-3H-indo]-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]-1-oxopropyl]-, nitrate (ester) (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

19

REFERENCE COUNT: THIS

THERE ARE 19 CITED REFERENCES AVAILABLE FOR

FORMAT

RECORD. ALL CITATIONS AVAILABLE IN THE RE

Page 36 SAEED

L4 ANSWER 39 OF 65
ACCESSION NUMBER:
DOCUMENT NUMBER:
139:390884
SUITCES
AUTHOR(S):
CORPORATE SOURCE:
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LANGUAGE: English
AB Receptor tyrosine kinase activation contributes to cell viability during
cytotoxic therapy. The novel broad spectrum receptor tyrosine kinase
inhibitor, SU1248, inhibits vascular endothelial growth factor receptor
2, platelet-derived growth factor receptor, c-kit, and fetal liver
tyrosine kinase 3. In this study, we maintained SU1248 plasma levels
beyond the completion of radiotherapy to determine whether tumor
regrowth can

the delayed. The antiangiogenic effects of SU11248 were demonstrated

using
human umbilical vein endothelial cells in vitro. Apoptosis increased and
clonogenic survival decreased when SU11248 was used in combination with
radiation from 0 to 6 Gy on endothelial cells. In vivo tumor growth

ction in vasculature was confirmed using the dorsal vascular window model. The vascular length established using images taken from a consistent quadrant in the window show the combination therapy was more effective in destroying tumor vasculature than either treatment alone. Sull248 maintenance administration beyond the completion of radiotherapy results in prolongation of tumor control. In summary, Sull248 enhances radiation-induced endothelial cytotoxicity, resulting in tumor vascular destruction and tumor control when combined with fractionated otherapy

destruction and tumor control when combined with fractionated radiotherapy
in murine tumor models. Moreover, inhibition of angiogenesis well beyond radiation therapy may be a promising treatment paradigm for refractory human neoplasms.

53795-19-4, SU11248
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological\_study); USES (Uses)
(SU11246 maintenance therapy prevents tumor regrowth after radiation)
RN 55795-19-4 CAPIUS
CN 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[{2}-(5-fluoro-1,2-dhydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (SCI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2003:551511 CAPLUS
DOCUMENT NUMBER: 139:101028
TITLE: Preparation of pyrrolylmethyle 139:101028
Preparation of pyrrolylmethyleneindolones as protein kinase inhibitors and antitumor agents
Griffin, John H.; Briesewitz, Roger; Wray, Jonathan

INVENTOR (S):

Theravance, Inc., USA PCT Int. Appl., 39 pp. CODEN: PIXXD2 Patent English PATENT ASSIGNEE(S): SOURCE:

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

|      | PAT           | TENT | NO.          |      |                 | KIN                                | D   | DATE |      |     | APPL | IÇAT | ION    | NO. |     |      | DATE  |     |
|------|---------------|------|--------------|------|-----------------|------------------------------------|-----|------|------|-----|------|------|--------|-----|-----|------|-------|-----|
|      | WO 2003057690 |      |              |      |                 |                                    |     |      |      |     |      |      |        |     |     |      |       |     |
|      |               |      |              |      | AM, AT, AU, AZ, |                                    |     |      |      |     |      |      |        |     |     |      |       |     |
|      |               |      |              |      |                 |                                    |     |      |      |     |      |      |        |     |     |      | GE,   |     |
|      |               |      |              |      |                 |                                    |     |      |      |     |      |      |        |     |     |      | LK,   |     |
|      |               |      |              |      |                 |                                    |     |      |      |     |      |      |        |     |     |      | OM,   |     |
|      |               |      |              |      |                 |                                    |     |      |      |     |      |      |        |     |     |      | TZ,   |     |
|      |               |      |              |      |                 |                                    |     |      |      |     |      | 10,  | ın,    | ın, | ıĸ, | 11,  | 12,   | UA, |
|      |               |      |              |      |                 |                                    |     | YU,  |      |     |      |      |        | 24  | 234 | 224  | 2.7   | DV  |
|      |               | KW:  | GH,          |      |                 |                                    |     |      |      |     |      |      |        |     |     |      |       |     |
|      |               |      |              |      |                 |                                    |     |      |      |     |      |      |        |     |     |      | EE,   |     |
|      |               |      |              |      |                 |                                    |     |      |      |     |      |      |        |     |     |      | BF,   | ы,  |
|      |               |      |              |      |                 |                                    |     | GN,  |      |     |      |      |        |     |     |      |       |     |
|      | US            | 2003 | 1713         | 78   |                 | A1 20030911 US 2002<br>B2 20040203 |     |      |      |     |      |      | 3273   | 85  |     | 2    | 20021 | 220 |
|      | US            | 6686 | 362          |      |                 | B2                                 |     | 2004 | 0203 |     |      |      |        |     |     |      |       |     |
|      | EΡ            |      | 713          |      |                 |                                    |     |      |      |     |      |      |        |     |     |      |       |     |
|      |               | R:   | AT,          |      |                 |                                    |     |      |      |     |      |      |        |     |     |      |       | ₽T, |
|      |               |      | IE,          | SI,  | LT,             | LV,                                | FI, | RO,  | ΜK,  | CY, | AL,  | TR,  | BG,    | CZ, | EE, | \$K  |       |     |
|      | BR            | 2002 | 0153<br>1988 | 60   |                 | A                                  |     | 2004 | 1214 |     | BR 2 | 002- | 1536   | 0   |     | 2    | 20021 | 220 |
| -    | US            | 2004 | 1988         | 04   |                 | Al                                 |     | 2004 | 1007 |     | US 2 | 003- | 6910   | 94  |     | 2    | 0031  | 022 |
| RIOR | IT            | APE  | LN.          | INFO | .:              |                                    |     |      |      |     | US 2 | 001- | 3437   | 46P |     | P 2  | 20011 | 227 |
|      |               |      |              |      |                 |                                    |     |      |      |     | US 2 | 001- | 3438   | 13P |     | P 2  | 0011  | 227 |
|      |               |      |              |      |                 |                                    |     |      |      |     | US 2 | 002- | 3273   | 85  |     | A3 2 | 0021  | 220 |
|      |               |      |              |      |                 |                                    |     |      |      |     | WO 2 | 002- | US 4 1 | 252 | ,   | w 2  | 0021  | 220 |

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

Title compds. I [wherein R1 = H, alkyl; R2 = -A1-NR5R6; R5, R6 = independently H, alkyl; A1 = (CH2)m, (CH2)m-A2-(CH2)p, (CH2CH2O)qCH2CH2;

= 2-10; n, p = 1-6; A2 = CH=CH, phenylene, biphenylene, cyclohexylene, piperazinylene; q = 1-3; or NR1R2 = morpholinyl, (un)substituted

azirinyl azirinyl, pyrrolidinyl, piperidinyl, azerown ethers, etc.;
R3, R4 = H, halo, alkyl, alkowy, unfaubstituted) Ph, SOZNHZ or alkyl/aryl
derivs., certain acylamino; and their pharmaceutically acceptable salts]
were prepared as receptor tyrosine kinase inhibitors useful in the

ANSWER 39 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

REFERENCE COUNT: THIS

THERE ARE 31 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

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ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) of proliferative disorders, such as cancer. For example, compd. II=(F3CCOZH)x was prepd. from 3,5-dimethyl-2,4-pyrrole dicarboxylic acid di-Et ester in Il steps via condensation with malonic acid, condensation with oxindole, amidation with mono-Boc piperaine and Boc deprotection using TFA. I exhibit an IC50 values of < 10 µM for inhibition of Fit-3, VEGFR and PDGFR tyrosinase kinases. II inhibited mutant Fit-3 tyrosinase kinase with EC50 = 0.24 µM.

560071-97-8P $60071-99-0P $60072-01-P9

560072-04-0P $60072-16-4P $60072-19-P9

560072-13-1P $60072-16-4P $60072-19-P9

560072-31-3P $60072-34-6P $60072-38-P9

560072-31-3P $60072-34-6P $60072-37-P9

S60072-31-3P $60072-34-FP $60072-37-P9

RI: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(Uses)
(Uses)
(tyrosine kinase inhibitor; preparation of
pyrrolylmethylenedihydroindolones
as protein kinase inhibitors and antitumor agents)
RN 560071-97-8 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-
N,2,4-trimethyl-N-[2-(methylamino)ethyl]-, mono(trifluoroacetate) (9CI)
(CA INDEX NAME)
                               CM 1
                               CRN 560071-96-7
CMF C22 H28 N4 O2
                                                                                                                                     O Me
|| |
| CH2-CH2-C-N-CH2-CH2-NHMe
                               CM 2
                                  CRN 76-05-1
CMF C2 H F3 O2
```

RN 560071-99-0 CAPLUS
CN 1H-Pyrcole-3-propanamide,
5-[(1,2-dhydro-2-oxo-3H-indol-3-ylidene)methyl]N,2,4-trimethyl-N-[3-(methylamino)propyl]-, trifluoroacetate (9CI) (CA
INDEX NAME)

CO2H

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L4 ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
                                                                                             (Continued)
         CH 1
        CRN 560071-98-9
CMF C23 H30 N4 O2
        CM 2
         CRN 76-05-1
CMF C2 H F3 O2
     -c-cо<sub>2</sub>н
 RN 560072-01-7 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-
N,2,4-trimethyl-N-[4-(methylamino)butyl]-, trifluoroacetate (9CI) (CA
NNEX NAME)
        CM 1
         CRN 560072-00-6
CMF C24 H32 N4 O2
        CM 2
        CRN 76-05-1
 L4 ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
F-C-CO2H
RN 560072-10-8 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-(11,2-dhydro-2-oxo-3H-indol-3-ylidene)methyl]-
N,2,4-trimethyl-N-[7-(methylamino)heptyl]-, trifluoroacetate (9CI) (CA INDEX NAME)
        CM 1
        CRN 560072-09-5
CMF C27 H38 N4 O2
```

```
L4 ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN CMF C2 H F3 O2
                                                                                                          (Continued)
 RN 560072-04-0 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-{(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-
N,2,4-trimethyl-N-{5-(methylamino)pentyl}-, trifluoroacetate (9CI) (CA INDEX NAME)
           CM 1
           CRN 560072-03-9
CMF C25 H34 N4 O2
          CM 2
           CRN 76-05-1
CMF C2 H F3 O2
 F-C-CO<sub>2</sub>H
 RN 560072-07-3 CAPLUS
CN 1H-Pyrcole-3-propanamide,
5-[(1,2-dh)ydro-2-oxo-3H-indol-3-ylidene)methyl]-
N,2,4-trimethyl-N-[6-(methylamino)hexyl}-, trifluoroacetate [9CI] (CA INDEX NAME)
          CM 1
          CRN 560072-06-2
CMF C26 H36 N4 O2
 L4 ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
                                                                                                          (Continued)
 RN 560072-13-1 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-
N,2,4-trimethyl-N-[8-(methylamino)octyl]-, trifluoroacetate (9CI) (CA INDEX NAME)
          CM 1
          CRN 560072-12-0
CMF C28 H40 N4 O2
          CM 2
          CRN 76-05-1
CMF C2 H F3 O2
F-C-CO2H
RN 560072-16-4 CAPLUS
CN 1H-Pyrcole-3-propanamide,
5-([1,2-dhydro-2-oxo-3H-indol-3-ylidene)methyl]-
N,2,4-trimethyl-N-[9-(methylamino)nonyl]-, trifluoroacetate (9CI) (CA
INDEX NAME)
          CM 1
          CRN 560072-15-3
CMF C29 H42 N4 O2
```

CM 2

L4 ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

CRN 76-05-1
CMF C2 H F3 O2

P—CO2H
F

RN 560072-19-7 CAPLUS
CN 1H-Pyrcole-3-propanamide,
5-[(1,2-dshydro-2-oxo-3H-indol-3-ylidene)methyl]N,2,4-trimethyl-N-[10-(methylamino)decyl]-, trifluoroacetate (9CI) (CA
INDEX NAME)

CM 1

CRN 560072-18-6
CMF C30 H44 N4 O2

CRN 76-05-1
CMF C2 H F3 O2

F—CO2H
F

RN 560072-22-2 CAPLUS
CN 1H-Pyrcole-3-propanamide,
5-[(1,2-dshydro-2-oxo-3H-indol-3-ylidene)methyl]N,2,4-trimethyl-N-[2-(methylamino)dodecyl]-, trifluoroacetate (9CI) (CA
INDEX NAME)

CM 1

L4 ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

F C CO2H

F

RN 560072-28-8 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-[(1,2-4hydro-2-oxo-3H-indol-3-ylidene)methyl)N,2,4-trimethyl-N-(2-[2-[2-(methylamino)ethoxy]ethoxy]ethyl]-,
trifluoroacetate (9CI) (CA INDEX NAME)

CM 1

CRN 560072-27-7
CMF C26 H36 N4 O4

O Me CH2-CH2-C-N-CH2-CH2-CH2-CH2-CH2-CH2-CH2-

PAGE 1-R

PAGE 1-A

L4 ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN CRN 560072-21-1 CMF C32 H48 N4 O2 (Continued) CM 2 CRN 76-05-1 CMF C2 H F3 O2 RN 560072-25-5 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-[(1,2-dthydro-2-oxo-3H-indol-3-ylidene)methyl]N,2,4-trimethyl-N-[4-(methylamino)-2-butenyl]-, trifluoroacetate (9CI)
(CA INDEX NAME) CM 1 CRN 560072-24-4 CMF C24 H30 N4 O2 CM 2 CRN 76-05-1 CMF C2 H F3 O2 ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) N-[2-[2-(2-ethoxyethoxy)ethoxy]-1-(methylamino)ethyl]-N, 2, 4-trimethyl-, trifluoroacetate (9CI) (CA INDEX NAME) CM 1 CRN 560072-30-2 CMF C28 H40 N4 O5 PAGE 1-A - CH2 - O - CH2 - CH2 - O - CH2 - -PAGE 1-B - CH2- OEt CM 2 CRN 76-05-1 CMF C2 H F3 O2 F- с- со<sub>2</sub>н RN 560072-34-6 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-[(1,2-dihydro-2-oxo-3H-indo]-3-ylidene)methyl]N,2,4-trimethyl-M-[[3-[(methylamino)methyl]phenyl]methyl]-,
trifluoroacetate (9CI) (CA INDEX NAME)

CRN 560072-33-5 CMF C28 H32 N4 O2 ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

CM 2

CRN 76-05-1 CMF C2 H F3 O2

RN 560072-37-9 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]N,2,4-trimethyl-M-[[4-{(methylamino)methyl)phenyl}methyl]-,
trifluoroacetate (9CI) (CA INDEX NAME)

CM 1

CRN 560072-36-8 CMF C28 H32 N4 O2

ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

CH 2

CRN 76-05-1 CMF C2 H F3 O2

RN 560072-43-7 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-

N, 2, 4-trimethyl-N-[{2'-{(methylamino)methyl][1,1'-biphenyl}-2-yl}methyl}-, trifluoroacetate (9CI) (CA INDEX NAME)

CM 1

CRN 560072-42-6 CMF C34 H36 N4 O2

ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

CM 2

CRN 76-05-1 CMF C2 H F3 O2

RN 560072-40-4 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]N,2,4-trimethyl-M-[[3-{(methylamino)methyl}cyclohexyl]methyl]-,
trifluoroacetate (9CI) (CA INDEX NAME)

CM 1

CRN 560072-39-1 CMF C28 H38 N4 O2

ANSWER 40 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

CM 2

CRN 76-05-1 CMF C2 H F3 O2

RN 560072-46-0 CAPLUS
CN 1H-Pyrrole-3-propanamide,
5-((1,2-dihydro-2-oxo-3H-indo1-3-ylidene)methyl]N,2,4-trimethyl-N-[3-[4-[3-(methylamino)propyl]-1-piperazinyl]propyl]-,
trifluoroacetate (9CI) (CA INDEX NAME)

CM 1

CRN 560072-45-9 CMF C30 H44 N6 O2

PAGE 1-A

W 20021218

MeNH- (CH2) 3

CM 2 76-05-1 C2 H F3 O2

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L4 ANSWER 42 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2003:532545 CAPLUS
DOCUMENT NUMBER: 139:95455
TITLE: Combined therapy against tumors comprising
substituted

acryloyl distamycin derivatives and protein kinase (serine/threonine kinase) inhibitors Geroni, Maria Cristina; Fowat, Camilla; Cozzi, Paolo Pharmacia Italia SpA, Italy PCT Int. Appl., 25 pp. CODEN: PIXXD2 Patent

INVENTOR(S): PATENT ASSIGNEE(S): SOURCE:

DOCUMENT TYPE: Patent

English 1

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO. APPLICATION NO. DATE KIND DATE 

WO 2002-EP13092

OTHER SOURCE(S):

MARPAT 139:95455

L4 ANSWER 41 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2003:534741 CAPLUS
DOCUMENT NUMBER: 139:214284
Early Amidation Approach to
3-[(4-Amido)pyrro1-2-y1]-2indolinones
AUTHOR(S): Manley, Jerad M.; Kalman, Monica J.; Conway, Brian G.:

AUTHOR(S):

Ball, Cynthia C.; Havens, Jeffrey L.; Vaidyanathan, Rajappa Chemical Research and Development, Pfizer Inc., Kalamazoo, MI, 49001, USA Journal of Organic Chemistry (2003), 68(16),

CORPORATE SOURCE:

cournal of Organic Chemistry (2003), 68(16),

6447-6450

CODEN: JOCEAH; ISSN: 0022-3263

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 139:214284

AB A new synthesis of 3-[(4-amido)pyrrol-2-yl]-2-indolinones has been developed, where the amide side chain was installed prior to pyrrole formation. This strategy precludes the need to use any coupling reagents to install the amide side chain. This process includes a zinc-free alternative to the Knorr pyrrole synthesis.

IT 55795-19-49

RL: SPN (Synthetic preparation): Page 1.

557795-19-4P
RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of 3-[(4-amido)pyrrol-2-yl]-2-indolinones via an early
amidation approach)
557795-19-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-((Z)-(5-fluoro-1, Z-dibydro-2-oxo-3H-indol-3-ylidene)methyl]-2, 4-dimethyl- (9CI) (CA INDEX
NAME)

Double bond geometry as shown.

REFERENCE COUNT: THIS

13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

ANSWER 42 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

The present invention provides the combined use of acryloyl distamycin derivs., in particular a-bromo- and a-chloro-acryloyl distamycin derivs., and a protein kinase (serine/threonine and tyrosine kinases) inhibitor, in the treatment of tumors. Also provided is the use of the said combinations in the treatment or prevention of metastasis or in the treatment of tumors by inhibition of angiogenesis. An example protein kinase inhibitor is STI 571 and a distamycin derivative is brostallicin (I).

557795-19-4

557795-19-4

(combined antitumor therapy comprising acryloyl distanycin derivs. and protein kinase (serine/threonine kinase) inhibitors)

557795-19-4

CAPLUS

1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl)-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown

REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT Page 41

SAEED

(Continued) ANSWER 43 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN NAME)

Double bond geometry as shown.

REFERENCE COUNT:

THERE ARE 78 CITED REFERENCES AVAILABLE FOR 78

FORMAT

RECORD. ALL CITATIONS AVAILABLE IN THE RE

L4 ANSWER 43 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN ACCESSION NUMBER: 2003:528824 CAPLUS DOCUMENT NUMBER: 140:70428 TITLE: Phase J despension 140:70428
Phase I dose-escalating study of SU11654, a small molecule receptor tyrosine kinase inhibitor, in dogs with spontaneous malignancies
London, Cheryl A.: Hannah, Alison L.: Zadovoskaya, Regina: Chien, May B.: Kollias-Baker, Cynthia;
Rosenberg, Mone: Downing, Sue: Post, Gerald; Boucher, Joseph: Shenoy, Narmada: Mendel, Dirk B.: McMahon, Gerald; Cherrington, Julie M.
School of Veterinary Medicine, University of California, Davis, CA, 95616, USA
Clinical Cancer Research (2003), 9(7), 2755-2768
CODEN: CCREF4; ISSN: 1078-0432
American Association for Cancer Research
Journal

AUTHOR (S):

SOURCE:

PUBLISHER:

TYPE: LANGUAGE:

CORPORATE SOURCE:

MENT TYPE: Journal JACE: English English The purpose of the following study was to investigate the safety and efficacy of the novel multitargeted indolinone receptor tyrosine kinase (RTK) inhibitor, SU1654, using a canine model of spontaneous tumors. This p.o. bloavailable compound exhibits potent inhibitory activity

This p.o. bloavailable compound exhibits potent inhibitory activity against members of the split kinase family of RTKs, including vascular endothelial growth factor receptor, platelet-derived growth factor receptor, Kit, and Flt-3, resulting in both direct antitumor and antiangiogenic activity. This was a Phase I trial in which successive cohorts of dogs with spontaneous tumors that had failed standard treatment regimens received escalating doses of SUIIG64 as oral therapy. Pharmacokinetics, toxicity, and tumor response were assessed. Fifty-seven dogs with a variety of cancers were enrolled; of these, 10 experienced progressive disease within

cancers were enrolled; of these, 10 experienced progressive disease within the first 3 Mk. Measurable objective responses were observed in 16 dogs (including 6 complete responses), primarily in mast cell tumors (n = 11), mixed mammary carcinomas (n = 2), soft tissue sarcomas (n = 2, and multiple myeloma (n = 1), for an overall response rate of 28 (16 of 57). Stable disease of sufficient duration to be considered clin. meaningful (>10 wk) was seen in an addnl. 15 dogs, for a resultant overall biol. activity of 548 (31 of 57). This study provides the first evidence that p.o. administered kinase inhibitors can exhibit activity against a variety

variety
of spontaneous malignancies. Given the similarities of canine and human
cancers with regard to tumor biol. and the presence of analogous RTK
dysregulation, it is likely that such agents will demonstrate comparable
antineoplastic activity in people.

IT 356069-94-5, SU11654
RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological
activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL
(Biological)

(Biological study); USES (Uses)

study); USES (Uses)
(small mol. receptor tyrosine kinase inhibitor SU11654 in dogs with spontaneous malignancies)
RN 356068-94-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[{Z}-{5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene}methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX

L4 ANSWER 44 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2003:491184 CAPLUS

DOCUMENT NUMBER: 193:69154 Preparation of 3-heteroarylmethylene-1,3-dihydro-2Hinvertor(s): Lin, Nah-Hering; Sham, Hing L.; Xia, Ping

Abbott Laboratories, USA

PCT Int. Appl., 21 pp.

CODENT TYPE: Patent

LANGUAGE: Patent

English

FAMILU ACC. NUM. COUNT: 1

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| PAT     | ENT  | NO.  |     |     | KIN | D   | DATE |      | AP    | PLI | CAT  | ION  | NO. |     | D.  | ATE  |     |
|---------|------|------|-----|-----|-----|-----|------|------|-------|-----|------|------|-----|-----|-----|------|-----|
|         |      |      |     |     |     | -   |      |      |       |     |      |      |     |     | -   |      |     |
| WO      | 2003 | 0518 | 38  |     | A2  |     | 2003 | 0626 | WO    | 20  | 02-1 | US39 | 641 |     | 2   | 0021 | 212 |
| WO      | 2003 | 0518 | 38  |     | A3  |     | 2003 | 0918 |       |     |      |      |     |     |     |      |     |
|         | W:   | CA,  | JP. | MX  |     |     |      |      |       |     |      |      |     |     |     |      |     |
|         | RW:  | AT,  | BE, | BG, | CH, | CY, | CZ,  | DE,  | DK, E | E,  | ES,  | FI.  | FR. | GB, | GR. | IE.  | IT. |
|         |      |      |     |     | PT, |     |      |      |       |     | -    |      |     |     |     |      |     |
| US      | 2003 | 1198 | 39  |     | Al  |     | 2003 | 0626 | US    | 20  | 01-  | 2229 | 0   |     | 2   | 0011 | 213 |
| EP      | 1453 | 800  |     |     | A2  |     | 2004 | 0908 | EP    | 20  | 02-  | 7900 | 89  |     | . 2 | 0021 | 212 |
|         | R:   | AT,  | BE, | CH, | DE, | DK, | ES,  | FR.  | GB, G | R,  | IT,  | LI.  | LU, | NL. | SE. | MC.  | PT. |
|         |      |      |     |     |     |     |      |      | EE, S |     |      |      |     |     | ,   | ,    |     |
| RIORITY | APP  |      |     |     |     |     |      |      |       |     | 01-  | 2229 | n   |     |     | 0011 | 212 |

WO 2002-US39641

OTHER SOURCE(S): MARPAT 139:69154

AB The title compds. [I; X = N, CR3; Y = N, CR4; Z = N, CR5; with the proviso

that at least one of Y and Z is other than N: one of R3-R5 and R1 = aryl or heterocyclyl and the others are H; R2 = aryl or heterocyclyl; with the proviso that when R2 is heterocyclyl, the heterocyclyl is other than imidazolyl] which are protein kinase inhibitors, were prepared Thus, reacting 6-bromo-1,3-dihydro-2H-indol-2-one with

1H-pyrrole-2-carbaldehyde in the presence of piperidine in MeOH followed by coupling of the resulting

(32)-6-bromo-3-(1H-pyrrol-2-ylmethylene)-1,3-dihydro-2H-indol-2-one with 4-tert-butyldimethylailyloxy-2-methylphenylboronic acid afforded (32)-1 [X, Y, Z = CH; R1 = 4-hydroxy-2-methylphenyl; R2 = pyrrol-2-yl]. The compds. I inhibited ChK1 at IC50 values between about 1 nM and about 10 µM.

10 μM. 550373-92-7P 550373-93-8P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

Page 42

SAEED

W 20021212

ANSWER 44 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) (Uses) (prepn. of 3-heteroarylmethylene-1,3-dihydro-2H-indol-2-ones as

protein

ein
 kinase inhibitors)
550373-92-7 CAPLUS
2H-Indol-2-one, 3-[[4-[(dimethylamino)methyl]-5-methyl-3-(1-methylethyl)1H-pyrrol-2-yl]methylene]-1, 3-dihydro-6-(4-hydroxy-3-methoxyphenyl)-,
(32)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

550373-93-8 CAPLUS
2N-Indol-2-one, 3-[[4-[(dimethylamino)methyl]-5-methyl-3-(1-methylethyl)-1R-pyrrol-2-yl]methylene]-1,3-dihydro-6-(4-hydroxy-3-methoxyphenyl)-,
(3Z)-, bis(trifluoroacetate) (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 550373-92-7 CMF C27 H31 N3 O3

Double bond geometry as shown.

CM 2

CRN 76-05-1 CMF C2 H F3 O2

L4 ANSWER 45 OF 65

ACCESSION NUMBER:

DOCUMENT NUMBER:

139:207284

SUIL284 inhibits KIT and platelet-derived growth factor receptor β in preclinical models of human small cell lung cancer

AUTHOR(S):

ADFAMS, Tinya J.; Lee, Leslie B.; Murray, Lesley J.; Pryer, Nancy K.; Cherrington, Julie M.

CORPORATE SOURCE:

SOURCE:

Molecular Cancer Therapeutics (2003), 2(5), 471-478 CODEN: MCTOCF; ISSN: 1535-7163

PUBLISHER:

DOCUMENT TYPE:

DOCUMENT TYPE:

DOCUMENT TYPE:

AB The purpose of this study was to evaluate the activity of the indolinone kinase inhibitor SUIL248 against the receptor tyrosine kinase KIT in

and in vivo, examine the role of KIT in small cell lung cancer (SCLC),

anticipate clin. utility of SU11248 in SCLC. SU11248 is an oral, multitargeted tyrosine kinase inhibitor with direct antitumor and antianglogenic activity through targeting platelet-derived growth factor receptor (PDGFR), vascular endothelial growth factor receptor, KTT, and FLT3 receptors. Treatment of the KIT-expressing SCLC-derived NCI-H526 cell line in vitro with SU11248 resulted in dose-dependent inhibition of stem cell factor-stimulated KIT phosphotyrosine levels and proliferation. The biol. significance of KIT inhibition was evaluated in vivo by ting

The biol. significance of the immediate by the SU11248 or another structurally unrelated KIT inhibitor, STI571 (Gleevec), which is also known to inhibit Ber-AbI and PDGFRB. SU11248 treatment resulted in significant tumor growth inhibition, whereas inhibition from STI571 treatment was less dramatic. Both compds. reduced phospho-KIT levels in NCI-H526 tumors, with a greater reduction by SU11248, correlating with efficacy. Likewise,

vise, phospho-PDGFRβ levels contributed by tumor stroma and with known involvement in angiogenesis were strongly inhibited by SU11248 and less

by STI571. Because platinum-based chemotherapy is part of the standard

care for SCLC, SU11248 was combined with cisplatin, and significant tumor growth delay was measured compared with either agent alone. These results

growth delay was measured compared with either agent alone. These
lts
expand the profile of SU11248 as a KIT signaling inhibitor and suggest
that SU11248 may have clin. potential in the treatment of SCLC via direct
antitumor activity mediated via KIT as well as tumor anglogenesis via
vascular endothelial growth factor receptor FLK1/KDR and PDGFRB.
ST795-19-4, SU11248
RL: DNA (Drug mechanism of action); PAC (Pharmacological activity); THU
(Therapeutic use); BIOL (Biological study); USES (Uses)
(SU11248 inhibits KIT and platelet-derived growth factor receptor
p in preclin. models of human small cell lung cancer)
ST795-19-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX
NAME)

Double bond geometry as shown.

L4 ANSWER 44 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

ANSWER 45 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

REFERENCE COUNT: THIS

32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L4 ANSWER 46 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2003:362945 CAPLUS
DOCUMENT NUMBER: 139:173373
TITLE: SU11248 is a novel FLT3 tyrosine kinase inhibitor

AUTHOR (5):

potent activity in vitro and in vivo
O'Farrell, Anne-Marie; Abrams, Tinya J.; Yuen, Helene
A.: Ngai, Theresa J.; Louie, Sharianne G.: Yee, Kevin
W. H.; Wong, Lily M.; Hong, Weiru; Lee, Leslie B.;
Town, Ajia; Smolich, Beverly D.; Manning, William C.;
Murray, Lesley J.; Heinrich, Michael C.; Cherrington,
Julie M.
Preclinical Research and Exploratory Development,
SUGEN, South San Francisco, CA, 94080, USA
Blood (2003), 101(9), 3597-3605
CODEN: BLOOW; ISSN: 0006-4971
American Society of Hematology
Journal

CORPORATE SOURCE:

SOURCE:

PUBLISHER:

PUBLISHER: American Society of Hematology
DOCUMENT TYPE: Journal
LANGUAGE: English
AB FLT3 (fms-related tyrosine kinase/flk2/Stk-2) is a receptor tyrosine
kinase (RTK) primarily expressed on hematopoietic cells. In blasts from
acute myelogenous leukemia (ANL) patients, 2 classes of FLT3 activating
mutations have been identified: internal tandem duplication (ITD)
mutations in the juxtamembrane domain (251-30% of patients) and point
mutations in the kinase domain activation loop (71-8% of patients).
FLT3-ITD mutations are the most common mol. defect identified in AML and
have been shown to be an independent prognostic factor for decreased
survival. FLT3-ITD is therefore an attractive mol. target for therapy.
SUI1248 is a recently described selective inhibitor with selectivity for
split kinase domain RTKs, including platelet-derived growth factor
receptors, vascular endothelial growth factor receptors, and KIT. We
show show

that SU11248 also has potent activity against wild-type FLT3 (FLT3-WT), FLT3-ITD, and FLT3 activation loop (FLT3-Asp835) mutants in phosphorylation assays. SU11248 inhibits FLT3-driven phosphorylation and induces apoptosis in vitro. In addition, SU11248 inhibits FLT3-induced VEGE

production The in vivo efficacy of SU11248 was investigated in 2 FLT3-ITD

models: a s.c. tumor xenograft model and a bone marrow engraftment model. We show that SU11248 (20 mg/kg/d) dramatically regresses FLT3-ITD tumors in the s.c. tumor xenograft model and prolongs survival in the bone

in the s.c. tumor xenograft model and prolongs survival in the bone ow engraftment model. Pharmacokinetic and pharmacodynamic anal. in s.c. tumors showed that a single administration of an efficacious drug dose potently inhibits FLT3-ITD phosphorylation for up to 16 h following a single dose. These results suggest that further exploration of SU11248 activity in AML patients is warranted.

S57795-19-4. SU11248
RE: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(FLT3 tyrosine kinase inhibitor SU11248 induces acute myelogenous leukemia apoptosis)
557795-19-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

L4 ANSWER 47 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN ACCESSION NUMBER: 2003:334853 CAPLUS DOCUMENT NUMBER: 138:331677 TITLE: Treatment of -----

138:331677
Treatment of acute myeloid leukemia with indolinone compounds, and preparation thereof O'Farrell, Ann-Marie; Cherrington, Julie Sugen, Inc., USA
PCT Int. Appl., 76 pp.
CODEN: PIXXD2
Patent INVENTOR (S)

PATENT ASSIGNEE (S): SOURCE:

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION: English

|               |      |    |  |      | KIND DATE |   |          |  |      |      |                      |     |     |     |      |      |
|---------------|------|----|--|------|-----------|---|----------|--|------|------|----------------------|-----|-----|-----|------|------|
|               |      |    |  |      |           | - |          |  |      |      |                      | -   |     |     |      |      |
| WO 2003035009 |      |    |  |      |           |   | 20030501 |  | WO 2 | 002- | 20021028             |     |     |     |      |      |
| WO            | 2003 | A3 |  | 2004 | 0318      |   |          |  |      |      |                      |     |     |     |      |      |
|               |      |    |  |      |           |   |          |  | BB,  | BG,  | BR,                  | BY, | BZ, | CA, | CH,  | CN   |
|               |      |    |  |      |           |   |          |  | EC,  |      |                      |     |     |     |      |      |
|               |      |    |  |      |           |   |          |  | KE.  |      |                      |     |     |     |      |      |
|               |      |    |  |      |           |   |          |  | MN.  |      |                      |     |     |     |      |      |
|               |      |    |  |      |           |   |          |  | SK,  |      |                      |     |     |     |      |      |
|               |      |    |  |      |           |   |          |  | ZM,  |      |                      |     |     | ,   | ,    |      |
|               | RW:  |    |  |      |           |   |          |  | SZ,  |      | UG.                  | ZM. | ZW. | AM. | A2.  | BY   |
|               |      |    |  |      |           |   |          |  | BG,  |      |                      |     |     |     |      |      |
|               |      |    |  |      |           |   |          |  | NL.  |      |                      |     |     |     |      |      |
|               |      |    |  |      |           |   |          |  | MR.  |      |                      |     |     | ,   | ,    |      |
| US            | 2003 |    |  |      |           |   |          |  |      |      |                      |     |     | 2   | 0021 | 028  |
|               |      |    |  |      |           |   |          |  |      |      | 20021028<br>20021028 |     |     |     |      |      |
|               |      |    |  |      |           |   |          |  | GR.  |      |                      |     |     |     |      |      |
|               | •••  |    |  |      |           |   |          |  | AL.  |      |                      |     |     |     |      |      |
| BR            | 2002 |    |  |      |           |   |          |  |      |      |                      |     |     |     |      | 02 R |
|               | APP  |    |  |      |           |   |          |  |      |      |                      |     |     |     |      |      |

WO 2002-US34525

W 20021028

OTHER SOURCE(S):

MARPAT 138:331677

(NR5) (CHR) pZ

A method of treating acute myeloid leukemia in patient pos. for FLT-3-ITD is described. The treatment is accomplished by administration of an indelinone compound (Markush included). Preparation of the compds. of

ANSWER 46 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN NAME) (Continued)

REFERENCE COUNT:

43 THERE ARE 43 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

ANSWER 47 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) invention, e.g. I, is described.
342641-94-SP 346405-32-1P 355068-97-87
452104-85-7P 452104-87-9P 452105-23-6P
452105-24-7P 515138-82-6P 557795-19-4P
RL: PAC (Fharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (indolinone derivative preparation for treatment of acute myeloid leukemia)
RN 342641-94-5 CAPLUS
RN 14-847-96-3 Carboxamide, N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

346405-32-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl|-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

356068-97-8 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(ethylamino)ethyl)-5-{(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

452104-85-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{Z}}-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

452104-87-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

452105-23-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[(ZS)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(SCI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

ANSWER 47 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

557795-19-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

499220-14-3
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (indolinone derivative preparation for treatment of acute myeloid

(indolinone derivative preparation for treatment of acute myeloid leukemia)
RN 499220-14-3 CAPLUS
CN Butanedioic acid, hydroxy-, (2S)-, compd. with
N-(2-(diethylamino)ethyl)-5(-(fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl-lH-pyrrole-3-carboxamide (1:1) (9CI) (CA INDEX NAME)

CM 1

Absolute stereochemistry. Rotation (-).

L4 ANSWER 47 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452105-24-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{Z}]-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-M-({ZR}-2-hydroxy-3-(4-morpholinyl)propyl)-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

515138-82-6 CAPLUS
IH-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-yildene)methyl]-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX

ANSWER 47 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

452104-42-6P 452104-86-8P 452104-88-0P 452104-89-1P 452104-90-4P 452104-91-5P 452104-92-6P 8E: SPN (Synthetic preparation); PREP (Preparation) (indolinone derivative preparation for treatment of acute myeloid

| leukemtal | RN | 452104-42-6 | CAPLUS | RN | 452104-42-6 | CAPLUS | RN | CAPLUS | CAPL

452104-86-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{2}-(1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl|-N-{2-hydroxy-3-(4-morpholinyl)propyl}-2,4-dimethyl- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

452104-88-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

452104-89-1 CAPLUS

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## 17/02/2005

10081147

ANSWER 47 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 1H-Pyrrole-3-carboxamide, 5-[(Z)-(1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-90-4 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl)-N-{2-hydroxy-3-(H-1,2,3-triazol-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-91-5 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl}-N-{2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

452104-92-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

L4 ANSWER 48 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2003:239162 CAPLUS
DOCUMENT NUMBER: 139:331991
TITLE: VEGF blocking therapy in the treatment of cancer
AUTHOR(S): Glade-Bender, Julia: Kandel, Jessica J.; Yamashiro,
Darrell J.
CORPORATE SOURCE: Divisions of Pediatric Oncology and Pediatric

CORPORATE SOURCE: Surgery,

College of Physicians and Surgeons at Columbia University, New York, NY, 10032, USA Expert Opinion on Biological Therapy (2003), 3(2), 263-276 CODEN: EOBTA2; ISSN: 1471-2598 Ashley Publications Ltd. Journal; General Review

SOURCE:

CODEN: EOBTA2: ISSN: 1471-2598

PUBLISHER: Ashley Publications Ltd.

DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

AB A review. It is widely accepted that tumor growth beyond a few cubic millimeters cannot occur without the induction of a new vascular supply. Inhibiting the development of new blood vessels (antianglogenesis) is a potential approach to cancer therapy that has attracted interest in recent

years. In theory, this approach should be relatively selective for tumor cells. The endothelial cells which form new vascular networks in tumors are responding to angiogenic stimuli produced by the tumor, but are themselves genetically normal. Endothelium in normal tissue, by

themselves genetically normal. Endothelium in normal tissue, by trast, themselves genetically normal. Endothelium in normal tissue, by trast, as usually quiescent. Vascular endothelial growth factor (VEGF) is the best-characterized pro-angiogenic factor. It is virtually ubiquitous in human tumnors, and higher levels have been correlated with more aggressive disease. Effective blockade of the VEGF pathway has been demonstrated with multiple agents: neutralizing antibody, receptor tyrosine kinase inhibitors, and ribozyme or antisense mols. targeting expression. Fromising preclin. data document the potential of these agents for tumor growth inhibition and even tumor regression, yet translation of novel therapeutics targeting the VEGF pathway to the clinic has proved a substantial challenge in itself. While showing clear evidence of antitumor activity over a broad spectrum of exptl. tumors, the proper selection, dose, timing and sequence of anti-VEGF treatment in human cancer is not at all obvious. Classic Phase I dose escalation trial design may need to be modified, as higher doses may not be optimal in all patients or for all tumors. In addition, alternate or secondary biol.

points (e.g., non-progression) may be needed for early phase studies to document true activity, so as not to abandon effective agents. Recent studies of the neutralizing antibody bevacizumab, and small mol. tyrosine kinase inhibitor SUS416, demonstrate that, while unlikely to be effective as monotherapy, incorporation of VEGF blockade into cytotoxic regimens

increase overall response rates. However, incorporation may also produce new toxicities, including thromboembolic complications and bleeding.

Newer oral agents, such as SU6668, SU11248, PTK/87/ZK222584 and ZD6474, are particularly interesting for their potential for chronic therapy. Future clin. trials are likely to build on past experience with stricter entry criteria, supportive care guidelines and the use of surrogate markers.

557795-19-4, SU11248

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(VEGF blocking therapy in treatment of cancer in relation to angiogenesis inhibition)

L4 ANSWER 47 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

ylidene)methyl)-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

ANSWER 48 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 557795-19-4 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as sh

REFERENCE COUNT:

THERE ARE 166 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE REFORMAT 166

Page 46 SAEED

### 17/02/2005

#### 10081147

L4 ANSWER 49 OF 65 CAPLUS COPYRIGHT 2005 ACS ON STN
ACCESSION NUMBER: 2003:170636 CAPLUS
DOCUMENT NUMBER: 138:37929
DIACOVERU CF 5 CF 7

138:337929
Diacovery of 5-[5-Fluoro-2-oxo-1,2-dihydroindol-(32]-ylidenemethyl]-2,4-dimethyl-1H-pyrrola-3-carboxylic Acid (2-Dlethylaminoethyl)amide, a Novel Tyrosine Kinase Inhibitor Targeting Vascular Endothelial and Platelet-Derived Growth Factor Receptor Tyrosine

AUTHOR (S):

Platelet-Derived Growth Factor Receptor Tyrosine Kinase Sun, Lir Liang, Chris; Shirazian, Sheri; Zhou, Yong; Miller, Todd; Cui, Jean; Fukuda, Juri Y.; Chu, Ji-Yu; Nematalla, Asaad; Wang, Xueyan; Chen, Hui; Sistla, Anand; Luu, Tony C.; Tang, Flora; Wei, James; Tang, Cho SUGEN Inc., South San Francisco, CA, 94080, USA Jöürnal of Medicinal Chemistry (2003), 46(7), 1116-1119

CORPORATE SOURCE: SOURCE:

CODEN: JMCMAR; ISSN: 0022-2623 American Chemical Society

PUBLISHER: DOCUMENT TYPE: LANGUAGE:

Journal

OTHER SOURCE(S):

English CASREACT 138:337929

AB To improve the antitumor properties and optimize the pharmaceutical properties including solubility and protein binding of indolin-2-ones, a series

os of different basic and weakly basic pyrrolylmethylidene indolinones I [Rl = H, F, Cl, Br: R2 = Et2NCH2CH2, pyridin-4-ylmethyl, 2-(1,2,3-triazol-1-yl)ethyl, etc.] were designed and synthesized. Indolinone I [Rl = F, R2

Et2NCH2CH2 (II)] showed the best overall profile in terms of potency for the VEGF-R2 and PDGF-Rβ tyrosine kinase at biochem, and cellular levels, solubility, protein binding, and bioavailability. II is sently in phase I clin. trials for the treatment of cancers. 142641-63-89 342641-63-98-9459 4605-32-19 356068-90-19 356068-91-29 356068-90-37 551338-83-87 515138-83-67 515138-83-79 515138-83-87 515138-83 515138-83 515138-83 515138-83 515138-83 515138-83 515138-83 515138-83 515138-83 515138-83 515138-83 515138-83 515138-83 515138-83 515138-83 515138-83 515138-

ANSWER 49 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

356068-90-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX

356068-91-2 CAPLUS
1H-Pyrrole-3-carboxamide,
(diethylamino)ethyl)-5-((1,2-dihydro-2-oxo3H-indol-3-ylidene)methyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

1H-Pyrrole-3-carboxamide, N-[2-[dimethylamino]ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

ANSWER 49 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RL: PAC (Pharmacological activity); SFN (Synthetic preparation); BIOL
(Biological study); PREP (Preparation)
(prepn. of (pyrron)ylmethylidene)indolinones as tyrosine kinase
inhibitors targeting vascular endothelial and platelet-derived growth
factor receptor tyrosine kinase)
342641-63-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (SCI) (CA INDEX
NAME)

342641-94-5 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

346405-32-1 CAPLUS

NH 374703-3-1 CAFLOS

NH-Pyrrole-3-cathoxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA

NAME)

ANSWER 49 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

515138-81-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[(1-methyl-4-piperidinyl)methyl]- (9CI)
(CA INDEX NAME)

515138-82-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAME)

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515138-83-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-(4-pyridinylmethyl)- (9CI) (CA INDEX

515138-84-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl)- (9CI)
(CA INDEX NAME)

L4 ANSWER 49 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

REFERENCE COUNT:

THERE ARE 13 CITED REFERENCES AVAILABLE FOR RECORD. ALL CITATIONS AVAILABLE IN THE RE

(Continued)

L4 ANSWER 50 OF 65
ACCESSION NUMBER:
DOCUMENT NUMBER:
138:193306
Preparation of crystals of a malic acid salt of
N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2oxo-3H-indole-3-ylidene)methyl]-2,4-dimethyl-1Hpyrrole-3-carboxamide
Hawley, Michael; Pieck, Thomas J.; Prescott, Stephen
P.; Maloney, Mark T.
PATENT ASSIGNEE(S):
SOURCE:
DOCUMENT TYPE:
LANGUAGE:
FAMILY ACC. NUM. COUNT:
1 English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| PA'      | TENT :        |       |      |     |     |     | DATE |      |     |       |       |      | DATE |     |          |      |     |  |  |
|----------|---------------|-------|------|-----|-----|-----|------|------|-----|-------|-------|------|------|-----|----------|------|-----|--|--|
| WO       | WO 2003016305 |       |      |     |     |     |      |      |     |       |       |      |      |     |          |      |     |  |  |
|          | W: AE, AG, Al |       |      | AL, | AM, | AT, | ΑU,  | AZ,  | BA, | BB,   | BG,   | BR,  | BY,  | BZ, | CA,      | CH,  | CN, |  |  |
|          |               | co,   | CR,  | CU, | CZ, | DE, | DK,  | DM,  | DZ, | EC,   | EE,   | ES,  | FI,  | GB, | GD,      | GE,  | GH, |  |  |
|          |               | GM,   | HR,  | HU, | ID, | IL, | IN,  | IS,  | JP, | KE,   | KG,   | KP,  | KR,  | KZ, | LC,      | LK,  | LR, |  |  |
|          |               | LS,   | LT,  | LU, | LV, | MA, | MD,  | MG,  | MK, | MN,   | MW,   | MX,  | MZ,  | NO, | NZ,      | OM,  | PH, |  |  |
|          |               | PL,   | PT,  | RO, | RU, | SD, | SE,  | SG,  | SI, | sĸ,   | SL,   | TJ,  | TM,  | TN, | TR,      | TT,  | TZ, |  |  |
|          |               | UΑ,   | UG,  | υs, | UΖ, | VC, | VN,  | ΥU,  | ZA, | ZM,   | ZW,   | AM,  | ΑZ,  | BY, | KG,      | ΚZ,  | MD, |  |  |
|          |               | RU,   | ΤJ,  | TM  |     |     |      |      |     |       |       |      |      |     |          |      |     |  |  |
|          | RW:           | GH,   | GM,  | ΚE, | LS, | MW, | ΜZ,  | SD,  | SL, | SZ,   | ΤZ,   | UG,  | ZM,  | ZW, | AT,      | BE,  | BG, |  |  |
|          |               | CH,   | CY,  | CZ, | DΕ, | DK, | EE,  | ES,  | FI, | FR,   | GB,   | GR,  | ΙE,  | IT, | LU,      | MC,  | NL, |  |  |
|          |               | PT,   | SE,  | SK, | TR, | BF, | ВJ,  | CF,  | CG, | CI,   | CM,   | GΑ,  | GN,  | GQ, | G₩,      | ML,  | MR, |  |  |
|          |               |       | SN,  |     |     |     |      |      |     |       |       |      |      |     |          |      |     |  |  |
|          |               |       |      |     |     |     |      |      |     |       |       |      |      |     | 20020813 |      |     |  |  |
| EP       |               |       |      |     |     |     |      |      |     |       |       |      |      |     | 20020813 |      |     |  |  |
|          | R:            |       |      |     |     |     | ES,  |      |     |       |       |      |      |     |          | MC,  | PT, |  |  |
|          |               |       |      |     |     |     | RO,  |      |     |       |       |      |      |     |          |      |     |  |  |
| BR       | 2002          | 0116  | 12   |     | A   |     | 2004 | 0824 |     | BR 21 | 002-  | 1161 | 2    |     | 2        | 0020 | B13 |  |  |
| NZ       | 5312          | 32    |      |     | A   |     | 2004 | 1126 | 1   | NZ 21 | 002-  | 5312 | 32   |     | 2        | 0020 | 813 |  |  |
| JP       | 2005          | 50331 | 86   |     | Т2  |     | 2005 | 0203 |     | JP 21 | 003-  | 5212 | 28   |     | 2        | 0020 | 813 |  |  |
| PRIORITY | APP           | LN.   | INFO | .:  |     |     |      |      | 1   | US 21 | 001-  | 3123 | 53P  |     | P 2      | 0010 | B15 |  |  |
|          |               |       |      |     |     |     |      |      | 1   | WO 2  | 002-1 | US25 | 649  | 1   | ¥ 2      | 0020 | 813 |  |  |

The present invention provides crystals, and compns. of the title

ound
Methods of preparing such crystals are also disclosed. Thus,
N-[2-(dicthylamino)ethyl]-5-[(5-fluoro-2-oxo-3H-indole-3-ylidene)methyl]2,4-dimethyl-1H-pyrrole-3-carboxamide. was dissolved in MeOH and L-malic
acid wass added to the solution The MeOh was removed and MeCN was added

and
the slurry was stirred and heated for 10 min. After cooling, the crystal form I of the title compound was obtained.

IT 499220-14-3P
RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PRP (Preparation); USES (Uses) (preparation of crystals of malic acid sait of (diethylamino)ethyl(fluorooxo indolyidene)methyldimethylpyrrolecarboxamide)
RN 499220-14-3 CAPLUS
CN Butanedioic acid, hydroxy-, (2S)-, compd. with N-[2-(diethylamino)ethyl]-5-

ANSWER 50 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) [(5-fluoro-1,2-dihydro-2-oxo-3H-lndol-3-ylidene)methyl]-2,4-dimethyl-lH-pyrrole-3-carboxamide (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 342641-94-5 CMF C22 H27 F N4 O2

CM 2

Absolute stereochemistry. Rotation (-).

IT 342641-94-5
RL: RCT (Reactant); THU (Therapeutic use); BIOL (Biological study); RACT (Reactant or reagent); USES (Uses) (Comparation of crystals of malic acid salt of (diethylaminolethyl(fluorooxo indolyldene)methyldimethylpyrrolecarboxamide)
RN 342641-94-5 CAPLUS
RN 342641-94-5 CAPLUS
RN 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

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L4 ANSWER 50 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

(Continued)

| MAIN | PATENT NO.

WO 2003015608
W1: AE, AG, AL,
CO, CR, CU,
GM, HR, HU,
LS, LT, LU,
-PL, PT, RO,
UA, UG, US,
RW: GH, GM, KE,
KG, KZ, MD,
FI, FR, GB,
CG, CI, CM,
US 2003216410
BR 2002011978
JP 2005501843
PRIORITY APPLN: INFO:: WO 2002-US25797 W 20020815 OTHER SOURCE(S): MARPAT 138:180703

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2003:154170 CAPLUS
DOCUMENT NUMBER: 138:180703
Combination therapy for the treatment of cancer
INVENTOR(8): Doshi, Parul; Cherrington, Julie
Masferrer, Jaine, USA; Sugen Inc.
FOR TITL Appl., 217 pp.
COODE: PIXXD2
DOCUMENT TYPE: PATENT INFORMATION: English
FAMILY ACC. NUM. COUNT: 1

DATE

KIND

APPLICATION NO.

DATE

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO.

AB The present invention relates to methods for treatment or prevention of neoplasia disorders using protein tyrosine kinase inhibitors in combination with cyclooxygenase inhibitors, in particular cyclooxygenase-2

ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 2,3-dihydro-lH-indole-4-carboxylic acid (3-chloro-4-methoxyphenyl) amide 342641-82-1P, 5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)- 2,4-dimethyl-1H-pyrrole-3-carboxylic acid (3-diethylaminopropyl) amide 342641-83-2P, 5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)- 2,4-diisporpoyl-1H-pyrrole-3-carboxylic acid (2-diethylaminocethyl) amide 342641-84-3P, 5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)- 2,4-dimethylaminopropyl-1H-pyrrole-3-carboxylic acid (3-diethylaminopropyl) amide 342641-85-4P 342641-87-6P, 5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (pyridin-4-ylmethyl)-amide 342641-88-8P,

5-{6-(5-Isopropyl-2-methoxyphenyl)-2-oxo-1,2-dihydroindol-3-ylidenemethyl}-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-pyrrolidin-1-ylethyl)amide 342641-91-2P 342641-92-3P, 5-{6-(2,4-Dimethoxyphenyl)-2-

342641-91-27 342641-92-37, 5-[6-[2,4-Dimethoxyphenyl]-2
oxo-1,2-dihydroindol-3-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic
acid (2-pyrrolidin-1-ylethyl)amide 342641-93-4P,
5-[6-(3-Isopropylphenyl]-2-oxo-1,2-dihydroindol-3-ylidenemethyl]-2,4dimethyl-1H-pyrrole-3-carboxylic acid (2-pyrrolidin-1-ylethyl)amide
342641-95-6F, 3-[4-(2-Diethylaminoethylcarbamoyl)-3,5-dimethyl-1Hpyrrol-2-ylmethylene]-2-oxo-2,3-dihydro-1H-indole-6-carboxylic acid
342641-96-7F, 5-(5-Dimethylsulfamoyl-2-oxo-1,2-dihydroindol-3ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
(2-pyrrolidin-1-ylethyl)amide 342641-97-8F 342641-98-9F
, 2,4-Dimethyl-5-[2-oxo-5-(pyridin-3-ylufamoyl)-1,2-dihydroindol-3ylidenemethyl]-1H-pyrrole-3-carboxylic acid (2-pyrrolidin-1-ylethyl)amide
342642-01-7F, 5-(5-Dimethylsulfamoyl-2-oxo-1,2-dihydroindol-3ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
(2-diethylaminoethyl)amide 342642-02-8F, 5-[5-(3-Chlorophenyl

ylideneentyl)-2,4-dimetryl-1n-pyrrole-3-carboxylic acid
(2-diethylaminoethyl) amide 342642-02-89, 5-[5-[3-Chlorophenyl)

sulfamoyl)-2-oxo-1,2-dihydroindol-3-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 342642-10-89
34262-11-99 356668-80-1P; 5-[5-Chloro-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-1-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-pyrrolidin-1-ylidenyl)-amide 355668-90-1P 356668-90-1P 356668-90-1P 356668-90-1P 356668-90-1P 356668-90-1P 356668-90-1P 35668-90-1P 35668-90-1

Double bond geometry as shown. ### 342641-49-0P 342641-50-3P 342641-51-4P 342641-32-5P 342641-50-3P 342641-55-8P 342641-55-9P 342641-55-9P 342641-55-9P 342641-59-9P 342641-59-9P 342641-65-9P 342641-65-9P 342641-65-9P 342641-65-9P 342641-65-9P 342641-65-9P 342641-67-PP 342641-67-PP 342641-70-PP 3

ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) selective inhibitors. The protein kinase inhibitors are of the formula I where R = H, piperazin-1-yimethyl, 4-methylpiperazin-1-yimethyl, piperidin-1-yimethyl, etc.; R1 = H, halo, alkyl, substituted alkyl, piperidin-1-yimethyl, etc.; R2 = H, halo, alkyl, substituted alkyl, substituted dycloalkyl, are, etc.; R3 = H, alogen, alkyl, substituted alkyl, trihalomethyl, hydroxy, alkoxy, etc.; R3 = H, halogen, alkyl, substituted alkyl, trihalomethyl, hydroxy, alkoxy, etc.; R4 = H, halogen, alkyl, substituted alkyl, hydroxy, alkoxy, etc.; R5 = H, alkyl, substituted alkyl, hydroxy, alkoxy, etc.; R5 = H, alkyl, substituted alkyl, alkoxy, etc.; R6 = H, alkyl, substituted alkyl, aryl, heteroaryl, etc.

186088-97-8P
R1: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); RTU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RTO (Reactant or reagent); USES (Uses) (combination therapy for treatment of cancer using protein tyrosine kinase inhibitors and cyclooxygenase-2 inhibitors)

186088-97-8 CAPLUS

18-Pyrrole-3-carboxamide, N-{2-(ethylamino)ethyl]-5-{(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

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# 17/02/2005

#### 10081147

Answer S1 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
356069-46-09 356069-47-1P 356069-48-2P
356069-49-3P 356069-50-6P 356069-51-7P
356069-30-3P 356069-55-1P 356069-57-7P
356069-58-4P 356069-55-1P 356069-60-8P
356069-61-0P 356069-62-0P 356069-64-2P
356069-65-3P 356069-66-4P 356069-77-7P
357798-1P-4P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
{combination therapy for treatment of cancer using protein tyrosine kinase inhibitors and cyclooxygenase-2 inhibitors}
342641-49-0 CAPLUS
HI-Pyrrole-3-carboxamide, 5-{(5-bromo-1, 2-dihydro-2-cxo-3H-indol-3-ylidene)methyl]-N-{3-(diethylamino)propyl]-2-(1-methylethyl)-4-phenyl-(9CI) (CA INDEX NAME)

342641-50-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylideno|methyl|-2-(1-methylethyl)-4-phenyl-N-[3-(1-pyrrolidinyl)propyl]-(9CI) [CA INDEX NAME)

342641-51-4 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

342641-55-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-(2-methoxyphenyl)-2-oxo-3H
indol-3-ylidene|methyl]-2-methyl-4-phenyl-N-[2-(1-pyrrolidinyl)ethyl](9CI) (CA INDEX NAME)

342641-56-9 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-[2-(dimethylamino)ethyl]-2-methyl-4-phenyl- (9CI) (CA INDEX NAME)

ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene)methyl]-N-[2-(diethylamino)ethyl]-2-(1-methylethyl)-4-phenyl-(SCI) (CA INDEX NAME)

$$\begin{array}{c} \text{Br} \\ \\ \text{Ph} \\ \\ \end{array} \begin{array}{c} \text{Ph} \\ \\ \text{Ph} \\ \end{array} \begin{array}{c} \text{Pr-i} \\ \\ \text{C-NH-CH}_2\text{-CH}_2\text{-NEt}_2 \\ \end{array}$$

342641-52-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2-(1-methylethyl)-N-[3-(4-methyl-1-piperazinyl)propyl]-4-phenyl- (9CI) (CA INDEX NAME)

342641-54-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2-methyl-4-phenyl-N-[2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX NAME)

ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

(Continued)

342641-57-0 CAPLUS

1H-Pyrrole-3-carboxamide, 5-{{1,2-dihydro-6-(2-methoxyphenyl)-2-oxo-3H-indol-3-ylidene)methyl}-N-[2-(dimethylamino)ethyl]-2-methyl-4-phenyl-(9CI) (CA INDEX NAME)

342641-59-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3ylidene)methyl]-N-[3-(diethylamino)propyl]-2-methyl-4-phenyl- (9CI) (CA
INDEX NAME)

342641-60-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-61-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl}-N-[2-(dimethylamino)ethyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-62-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-63-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

(Continued)

RN 342641-64-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl}- (9CI) (CA INDEX NAME)

RN 342641-65-0 CAPLUS
CN lH-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl)-N-[3-(lH-imidazol-1-yl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-66-1 CAPLUS
CN lH-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-(2-methoxyphenyl)-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-67-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-(3-methoxyphenyl)-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI)
(CA INDEX NAWE)

RN 342641-68-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(dicthylamino)ethyl]-5-[(1,2-dihydro-2-oxo5-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-69-4 CAPLUS CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl)-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX NDEX)

RN 342641-70-7 CAPLUS
CN IH-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl]-N-[3-(1H-imidazol-1-yl)propyl]-2, 4-dimethyl- (9CI) (CA INDEX NAME)

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RN 342641-71-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-72-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-75-2 CAPLUS
CN 1H-Pytrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-[[1,2-dihydro-2-oxo-6-(3-pyridinyl)-3H-indol-3-ylidene]methyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

RN 342641-76-3 CAPLUS CN 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-6-(3-pyridinyl)-3H-indol-3ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME) L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-73-0 CAPLUS

(N lH-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-N-[3-(1H-imidazol-1-yl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-74-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[6-(3,5-dichlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene|methyl]-N-[2-{diethylamino}ethyl]-2,4-dimethyl- (9CI)
(CA
INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued

RN 342641-77-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[1,2-dihydro-2-oxo-6-[3-pyridinyl)-3H-indol-3ylidene]methyl]-N-[3-(dimethylamino)propyl]-2,4-dimethyl- (9CI) (CA
INDEX

RN 342641-78-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene|methyl]-N-[3-(dimethylamino)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

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L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 342641-79-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[3-(diethylaminolpropyl]-5-[(1,2-dihydro-2-oxo5-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-80-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-{3-{dichylaminolyropyl}-5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

342641-81-0 CAPLUS
1H-Indole-4-carboxamide, N-(3-chloro-4-methoxyphenyl)-3-[[4-[[(3-(diethylamino]propyl]amino]carbonyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo- (9CI) (CA INDEX NAME)

ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (CA INDEX NAME) (Continued)

RN 342641-85-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-bis(1-methylethyl)-N-[3-(1-pyrrolidinyl)propyl]-(9CI)

(CA INDEX NAME)

RN 342641-87-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-yildene)methyl]-2,4-dimethyl-N-(4-pyridinylmethyl)- (9CI) (CA INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-82-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[3-(diethylamino)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

342641-83-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-bis(1-methylethyl)- (9CI)(CA INDEX NAME)

342641-84-3 CAPLUS
1H-Pyrcole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3ylidene)methyl]-N-[3-(diethylamino)propyl]-2,4-bis(1-methylethyl)- (9CI)

ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

342641-88-7 CAPLUS
1H-Pyrrole-3-carboxamide,
[6-(4-butylphenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene|methyl|-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

342641-89-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-[2-methoxy-5-(1-

methylethyl)phenyl}-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX NAME)

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OME H O H ME O NH CH2 CH2

RN 342641-91-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[6-(4-ethylphenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene|methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Et H O H Me O NH CH2 CH2

RN 342641-92-3 CAPLUS
CN 1H-Pytrole-3-carboxamide,
5-[[6-(2,4-dimethoxyphenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pytrolidinyl)ethyl]- (9CI)
(CA INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued

RN 342641-96-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[[5-[(dimethylamino)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]-(9C1) (CA INDEX NAME)

RN 342641-97-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[[5-[(3-chlorophenyl)amino]sulfonyl]-1,2-dinydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-93-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-[3-(1-methylethyl)phenyl]-2-oxo-3M-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]-(9CI) (CA INDEX NAME)

RN 342641-95-6 CAPLUS
CN 1H-Indole-6-carboxylic acid,
3-[4-[[2-(diethylamino)ethyl]amino]carbonyl
|-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo- (9CI) (CA
INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued

RN 342641-98-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[{1,2-dihydro-2-oxo-5-[(3-pyridinylamino)sulfonyl]-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)dethyl]- (9CI) (CA INDEX NAME)

RN 342642-01-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[[5-

{ (dimethylamino) sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

Me2N-S CH NH-CH2-CH2-NEt2

RN 342642-02-8 CAPLUS

RN H-Pyrrole-3-carboxamide, 5-[[5-[[(3-chlorophenyl)amino]sulfonyl]-1,2-dihydro-2-oxo-34-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342642-10-8 CAPLUS
CN Glycine, N-[[5-[1,2-dihydro-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-ylidenejmethyl]-4-methyl-1H-pyrrol-3-yl]carbonyl]-, ethyl ester (9CI)

(CA INDEX NAME)

MeNH-S CH-CH2-S OEt

RN 342642-11-9 CAPLUS
CN Glycine, N-[[5-[[1,2-dihydro-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-ylidene|methyl]-4-methyl-1H-pyrrol-3-yl]carbonyl}- (9CI) (CA INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 356068-91-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino) ethyl]-5-[(1,2-dihydro-2-oxo3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 356068-92-3 CAPLUS CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX NAME)

H Me

CH Me

CH2

CH2

NN

RN 356068-95-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

MeNH-S CH H CH2-CO2H

RN 356068-82-1 CAPLUS
CN 1H-Fyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX

C1 H Ne Ne Ne NH NH CH2 CH2

RN 356068-90-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-{2-(diethylamino)ethyl}-2,4-dimethyl- {9CI} (CA INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

H Me CH — CH2 — CH2 — NMe2

RN 356068-96-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide, N-{2-(dimethylamino)ethyl}-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

HH ME

CH HH ME

NH-CH2-CH2-NMe2

RN 356068-99-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethyloxidoamino)ethyl]-5-[(Z)-(5-fluoro1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

H O Me N Et

RN 356069-03-9 CAPLUS
CN IH-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2, 4-dimethyl-N-[2-(4-methyl-1-piperazinyl)ethyl]- (9CI) (CA INDEX NAME)

356069-04-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-{2-(4-methyl-1-piperazinyl)ethyl}- (9CI)
(CA INDEX NAME)

356069-05-1 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

356069-09-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{Z}-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(3,5-dimethyl-1-piperazinyl)ethyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-12-0 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(3,5-dimethyl-1-piperazinyl)ethyl]-5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-(9CI)

(CA INDEX NAME)

Double bond geometry as shown.

356069-13-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{{Z}}-{5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-{2-{3,5-dimethyl-1-piperazinyl}ethyl}-2,4-dimethyl-{SCI} (CA INDEX NAME)

ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene|methyl]-2,4-dimethyl-N-{2-(4-methyl-1-piperazinyl)ethyl}- (9CI) (CA INDEX NAME)

RN 356069-07-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(1,2-4h)ydro-2-oxo-3H-indol-3-ylidene)methyl]2,4-dimethyl-N-[2-(4-methyl-1-piperarinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN Double bond geometry as shown. (Continued)

356069-15-3 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-(2-(3,5-dimethyl-1-piperazinyl)ethyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

355069-24-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(acetylamino)ethyl]-5-[(Z)-(5-chloro-1,2-dhydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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356069-25-5 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(acetylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-26-6 CAPLUS 1H-Pyrcle-3-carboxamide, N-{2-(acetylamino)ethyl}-5-[(2]-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (SCI) (CA INDEX NAME)

Double bond geometry as shown.

356069-36-8 CAPLUS lH-Pyrrole-3-carboxamide,  $5-\{(2)-\{5-cyano-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl\}-2,4-dimethyl-N-\{3-\{2-oxo-1-pyrrolidinyl\}propyl\}-\{CA INDEX NAME)$ 

Double bond geometry as shown.

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356069-39-1 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(2-oxo-1-imidazolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-40-4 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(z)-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(2-oxo-1-imidazolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

356069-37-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{Z}]-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(2-oxo-1-imidazolidinyl)ethyl]- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

356069-38-0 CAPLUS lH-Pyrrole-3-carboxamide,  $5-\{(z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene]$  whethyl]-2,4-dimethyl-N- $\{2-(2-oxo-1-imidazolidinyl)ethyl\}-$  (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

356069-41-5 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(2)-(5-cyano-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(2-oxo-1-imidazolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-42-6 CAPLUS 1-Piperazineacetic acid, 4-[2-[{[5-[{Z}]-(5-bromo-1,2-dihydro-2-oxo-3H-

indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]amino]ethyl], ethyl ester (9CI) (CA INDEX NAME)

356069-43-7 CAPLUS
1-Piperazineacetic acid, 4-[2-[[[5-{(Z)-(5-chloro-1,2-dihydro-2-oxo-3Hindol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl}amino]ethyl], ethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-44-8 CAPLUS
1-Piperazineacetic acid, 4-[2-[[5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-

indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]amino]ethyl], ethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene)methyl]-N-[3-(hexahydro-2-oxo-1H-azepin-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

356069-48-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[3-(hexahydro-2-oxo-1H-azepin-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

356069-49-3 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[3-(hexahydro-2-oxo-1H-azepin-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

356069-50-6 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[{2}-(5-cyano-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl)-N-[3-(hexahydro-2-oxo-1H-azepin-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

356069-45-9 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-[(cyanomethyl)amino]ethyl]-5-[(2)-(1,2-dlhydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-46-0 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(2)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl)-N-{3-(hexahydro-2-oxo-1H-azepin-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-47-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-((Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356069-51-7 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(acetylamino)ethyl]-5-{(2)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-53-9 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-{2-(3-oxo-1-piperazinyl)ethyl}-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CRN 356069-52-8 CMF C22 H24 F N5 O3

356069-55-1 CAPLUS lH-Pyrrole-3-carboxamide, 5-[(Z)-(1,2-dihydro-2-oxo-3H-indol-3-ylidene) methyll-2,4-dimethyll-N-[2-(3-oxo-1-piperazinyl)ethyl)-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

Double bond geometry as shown.

CM 2

ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356069-59-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-(2-[(2-cyanoethyl)amino]ethyl]-2,4-dimethyl-(9CI) (CAINDEX NAME)

Double bond geometry as shown.

356069-60-8 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-[(2-cyanoethyl)amino]ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

 $356069-61-9 \quad CAPLUS \\ 1H-Pyrrole-3-carboxamide, \quad N-\{2-\{(2-cyanoethyl) amino\}ethyl\}-5-\{\{Z\}-(1,2-dhydro-2-oxo-3H-indol-3-ylidene)methyl\}-2, \\ 4-dimethyl-\{9CI\} \quad (CA \ INDEX \ NAME)$ 

Double bond geometry as shown.

ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 356069-57-3 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-cyano-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(3-oxo-1-piperazinyl)ethyl]-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CH 1

CRN 356069-56-2 CMF C23 H24 N6 O3

Double bond geometry as shown.

CM 2 CRN 76-05-1 CMF C2 H F3 O2

Double bond geometry as shown.

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

356069-62-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-cyano-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-[(2-cyanoethyl)amino]ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

 $\begin{array}{lll} 356069-64-2 & CAPLUS \\ 1H-Pytrole-3-carboxamide, & 5-\{(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2, & 4-dimethyl-N-\{2-(3-oxo-1-piperazinyl)ethyl]-, \\ mono(trifluoroacetate) & (9CI) & (CA & INDEX NAME) \\ \end{array}$ 

CRN 356069-63-1 CMF C22 H24 C1 N5 O3

CRN 76-05-1 CMF C2 H F3 O2

356069-65-3 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-(2-(4-methyl-1-piperazinyl)ethyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-66-4 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(2)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-{2-(4-methyl-1-piperazinyl)ethyl}- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

342641-94-5 346405-32-1 356069-16-6
356069-17-5
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
(Blological study); USES (Uses)
(combination therapy for treatment of cancer using protein tyrosine kinase inhibitors and cyclooxygenase-2 inhibitors)
342641-94-5 CAPLUS
1H-Pytrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

346405-32-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN {Continued}

356069-77-7 CAPLUS

350005-77-7 CAPLUS
1-Piperazineacetic acid, 4-[2-[[[5-[(2)-[1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]amino]ethyl]-, ethyl eater [9C1] (CA INDEX NAME)

Double bond geometry as shown.

557795-19-4 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-{diethylamino}ethyl]-5-{(Z)-{5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene}methyl}-2,4-dimethyl-{9CI} (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 356069-16-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(ethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 356069-17-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N(2-(dichyloxidoamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NARE)

342642-09-5
RL: RCT (Reactant); RACT (Reactant or reagent)
(combination therapy for treatment of cancer using protein tyrosine

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ANSWER 51 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) kinase inhibitors and cyclooxygenase-2 inhibitors) 342642-09-5 CAPLUS Glycine, N-{[5-[[1,2-dihydro-4-methyl-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-ylidene|methyl]-4-methyl-1H-pyrrol-3-yl]carbonyl]-, ethyl ester (9CI) (CA INDEX NAME)

NAME)

Double bond geometry as shown.

ANSWER 52 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) in human trials.

557795-19-4, SU11248
RL: DMA (Drug mechanism of action); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (in vivo antitumor activity of SU11248, a novel tyrosine kinase inhibitor targeting vascular endothelial growth factor and platelet-derived growth factor receptors: determination of a pharmacokinetic/pharmacodynamic relationship)

557795-19-4 CAPLUS
IH-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown

REFERENCE COUNT:

THERE ARE 27 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

L4 ANSWER 52 OF 65
ACCESSION NUMBER:
DOCUMENT NUMBER:
139:239801
In Vivo Antitumor Activity of SUll248, a Novel
Tyrosine Kinase Inhibitor Targeting Vascular
Endothelial Growth Factor and Platelet-derived Growth
Factor Receptors: Determination of a
Pharmacokinetic/Pharmacodynamic Relationship
Mendel, Dirk B.; Laird, A. Douglas; Xin, Xiaohua;
Louie, Sharianne G.; Christensen, James G.; Li,
Guangmin; Schreck, Randall E.; Abrams, Tinya J.;
Ngai,

Theresa J.; Lee, Leslie B.; Murray, Lesley J.;

Jeremy: Chan, Emily; Moss, Katherine G.; Haznedar, Joshua O.; Sukbuntherng, Juthamas: Blake, Robert A.; Sun, Li; Tang, Cho: Miller, Todd; Shirazian, Sheri; McMahon, Gerald; Cherrington, Julie M. Preclinical Research and Exploratory Development, SUGEN, Inc., South San Francisco, CA, 94080, USA Clinical Cancer Research (2003), 9(1), 327-337 CODEN: CCREF4; ISSN: 1078-0432 American Association for Cancer Research Journal English

CORPORATE SOURCE:

CODEN: CCREF4; ISSN: 1078-0432

PUBLISHER: American Association for Cancer Research
DOCUMENT TYPE: Journal
LANGUAGE: English

AB One challenging aspect in the clin. development of molecularly targeted
therapies, which represent a new and promising approach to treating
cancers, has been the identification of a biol. active dose rather than a
maximum tolerated dose. The goal of the present study was to identify a
pharmacokinetic/pharmacodynamic relationship in preclin. models that could

be used to help guide selection of a clin. dose. SU11248, a novel small mol. receptor tyrosine kinase inhibitor with direct antitumor as well as antiangiogenic activity via targeting the vascular endothelial growth factor (VEGF), platelet-derived growth factor (FDGF), KTT, and FLT3 receptor tyrosine kinases, was used as the pharmacol. agent in these studies. In mouse xenograft models, SU11248 exhibited broad and potent antitumor activity causing regression, growth arrest, or substantially reduced growth of various established xenografts derived from human or

tumor cell lines. To predict the target SU11248 exposure required to achieve antitumor activity in mouse xenograft models, we directly

measured
target phosphorylation in tumor xenografts before and after SU11248
treatment and correlated this with plasma inhibitor levels. In target
modulation studies in vivo, SU11248 selectively inhibited Flk-1/KDR (VEGF
receptor 2) and PDGF receptor B phosphorylation (in a time- and
dose-dependent manner; when plasma concra. of inhibitor reached or
exceeded 50-100 ng/ml. Similar results were obtained in a functional
assay of VEGF-induced vascular permeability in vivo. Constant
inhibition of

bition of VEGFR2 and PDGF receptor  $\beta$  phosphorylation was not required for efficacy; at highly efficacious doses, inhibition was sustained for 12 h of a 24-h dosing interval. The pharmacokinetic/pharmacokynamic relationship established for SU1248 in these preclin. studies has aided in the design, selection, and evaluation of dosing regimens being tested

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2002:927188 CAPLUS
DOCUMENT NUMBER: 138:14005
TITLE: Preparation of 5-aralkylsulfonyl-3-(pyrrol-2-ylmethylidene)-2-indolinone derivatives as kinase

inhibitors Innibicors Cui, Jingrong; Ramphal, Yudhi; Liang, Congxin; Sun, Li; Wei, Chung Chen; Tang, Peng Cho INVENTOR (S):

USA
PCT Int. Appl., 479 pp.
CODEN: PIXXD2
Patent PATENT ASSIGNEE(S): SOURCE:

DOCUMENT TYPE: English

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

|         | TENT          |      |      | KIND DATE |     |             |      |      | ICAT | DATE |      |      |     |     |     |      |     |
|---------|---------------|------|------|-----------|-----|-------------|------|------|------|------|------|------|-----|-----|-----|------|-----|
| WO      | 2002096361    |      |      |           |     | A2 20021205 |      |      |      |      | 002- |      |     |     |     |      |     |
| WO      | WO 2002096361 |      |      | A3        |     | 20030313    |      |      |      |      |      |      |     |     |     |      |     |
|         | W:            | ΑE,  | AG,  | AL,       | ΑM, | AT,         | AU,  | AZ,  | BA,  | BB,  | BG,  | BR,  | BY, | BZ, | CA, | CH,  | CN, |
|         |               | co,  | CR,  | Cυ,       | CZ, | DE,         | DK,  | DM,  | DZ,  | EC,  | EE,  | ES,  | FI, | GB, | GD, | GE,  | GH, |
|         |               | GM,  | HR,  | ΗU,       | ID, | IL,         | IN,  | IS,  | JP,  | KE,  | KG,  | KP,  | KR, | KZ, | LC, | LK,  | LR, |
|         |               | LS,  | LT,  | LU,       | LV, | MA,         | MD,  | MG,  | MK,  | MN,  | MW,  | MX,  | MZ, | NO, | NZ, | OM,  | PH, |
|         |               | PL,  | PT,  | RO,       | RU, | SD,         | SE,  | SG,  | SI,  | SK,  | SL,  | TJ,  | TM, | TN, | TR, | TT.  | TZ, |
|         |               | UA,  | UG,  | US,       | UZ, | VN,         | YU,  | ZA,  | ZM,  | ZW,  | AM,  | AZ,  | BY, | KG, | KZ, | MD.  | RU, |
|         |               | ТJ,  | TM   |           |     |             |      |      |      |      |      |      |     |     |     |      |     |
|         | RW:           | GH,  | GM,  | KE,       | LS, | MW,         | MZ,  | SD,  | SL,  | SZ,  | TZ,  | UG,  | ZM, | ZW, | AT, | BE,  | CH, |
|         |               | CY,  | DE,  | DK,       | ES, | FI,         | FR,  | GB,  | GR,  | IE,  | IT,  | LU,  | MC, | NL, | PT. | SE,  | TR, |
|         |               | ΒF,  | ВJ,  | CF,       | CG, | CI,         | CM,  | GΑ,  | GN,  | GQ,  | G₩,  | ML,  | MR, | NE, | SN, | TD,  | TG  |
| US      | 2003          | 1253 | 70   |           | A1  |             | 2003 | 0703 | 1    | US 2 | 002- | 1570 | 07  |     | 2   | 0020 | 530 |
| US      | 6599          | 902  |      |           | B2  |             | 2003 | 0729 |      |      |      |      |     |     |     |      |     |
| PRIORIT | Y APP         | LN.  | INFO | .:        |     |             |      |      | 1    | US 2 | 001- | 2945 | 44P |     | P 2 | 0010 | 530 |
|         |               |      |      |           |     |             |      |      | 1    | US 2 | 001- | 3284 | 08P |     | P 2 | 0011 | 010 |

OTHER SOURCE(S): MARPAT 138:14005

The present invention relates to certain 5-aralkylsulfonyl-3-(pyrrol-2-ylmsthylidene)-2-indolinone deriva. (abown as 1; see below for variable definitions; e.g. 2,4-dimethyl-5-(2-oxo-5-phenylmethanesulfonyl-1,2-

Page 61 SAEED ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) dihydroindol-(3Z)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)aminde) that inhibit kinases (no data), in particular met kinase. Pharmaceutical compns. comprising these compda., methods of treating diseases mediated by kinases using pharmaceutical compns. comprising these compda., and methods of prepg. them are also disclosed. In I: n = 0-2; m = 1-3; Rl and R2 = H or alkyl; R3, R4, and R5 = H, halo, alkyl, cycloalkyl, haloalkyl, hydroxy, alkoxy, alkoxycarbonyl, alkoxy.

alkyl, cycloalkyl, haloalkyl, hydroxy, alkoxy, alkoxycarbonyl, haloalkoxy, cyano, carboxy, carboxyalkyl, nitro, aryl, aryloxy, heteroaryl, heteroaryloxy, -[alkylene]-CONRIOR1, -CONRIOR1, or - NRIORII (R10 is H or alkyl, and R11 is aryl, heteroaryl, heterocycle, aminoalkyl, alkylaminoalkyl, dialkylaminoalkyl, dyndroxyalkyl, acetylalkyl, cyanoalkyl

alkylaminoalkyl, dialkylaminoalkyl, nyutosyaary, notymoalkyl, carboxyalkyl, alkoxycarbonylalkyl, heteroaralkyl, aralkyl, or heterocyclylalkyl wherein the alkyl chain in aminoalkyl, alkylaminoalkyl, dialkylaminoalkyl, aralkyl, heteroaralkyl, or heterocyclylalkyl is optionally substituted with one or two hydroxy, or R10 and R11 together with the N atom to which they are attached combine to form satd. or unsatd. heterocycloamino). R6 is H, alkyl, cycloalkyl, hydroxyalkyl, aminoalkyl, alkylaminoalkyl, dialkylaminoalkyl, carboxyalkyl, heterocyclylalkyl, aryl, heteroaryl, carboxy, alkoxycarbonyl, heterocyclylalkyl, aryl, heteroaryl, carboxy, alkoxycarbonyl, dialkylaminoalkylcarbonyl, dialkylaminoalkylcarbonyl, dialkylaminoalkylcarbonyl, dialkylaminoalkylcarbonyl, conR10R11 or -(alkylene)-CORR10R11. R7 and

H, alkyl, cycloalkyl, heterocyclylalkyl, -COR12, -(alkylene)-COR12 (R12 = alkoxy, hydroxy, or heterocycle, alkylamino, dialkylamino), -SO2R14, -CONR13R14, or -(alkylene)-CORR13R14 (R13 is H or alkyl, and R14 is aminoalkyl, alkylaminoalkyl, dialkylaminoalkyl, hydroxyalkyl,

aminoalryi, anykaminoagy, actylalkyl, actylalkyl, carboxyalkyl, cyanoalkyl, carboxyalkyl, alkoxycarbonylalkyl, heteroaralkyl, or heterocyclylalkyl wherein the alkyl chain in aminoalkyl, heteroaralkyl, heteroaralkyl, or heterocyclylalkyl is optionally substituted with one or two hydroxy group(s), or when R13 and R14 are attached to a N atom R13

R14 together with the N atom to which they are attached form satd. or unsatd. heterocycloamino). R6 and R7 or R7 and R8 can combine to form a satd. or unsatd. 5 to 8 membered ring; and R9 is: H or alkyl: -PO(OR15)2 where each R15 = H or alkyl: or -CHR17NR18R19 where R17 is H or alkyl, and R18 and R19 = H or alkyl or

other disorders) 477576-36-6 CAPLUS

ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-[1,2,3]triazol-1-ylethyl)amide 477573-80-1F,

2,4-Dimethyl-5-(4-methyl-2-oxo-5-phenylmethanesulfonyl-1,2-dihydroindol-3-(2)-ylidenemethyl)-H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477573-61-2P, 5-[5-(2-Fluorophenylmethanesulfonyl)-4-methyl-2-oxo-

1,2-dihydroindol-3-(Z)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477573-82-3P, 5-(2-chlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477573-83-4P, 4-{[[3-{1-[4-(2-

Diethylaminoethylcarbamoyl)-3,5-dimethyl-1H-pyrrol-2-yl]meth-[Z]-ylidene]2-oxo-2,3-dihydro-1H-indol-5-yl]sulfonyl]methyl]benzoic acid methyl ester
477573-84-5P, 5-[5-(4-Trifluoromethoxyphenylmethanesulfonyl)-2-oxo-

1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 477573-86-7P,
5-[5-[2,4-Bis (trifluoromethyl)] phenylmethanesulfonyl]-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 477573-88-9P,
5-[5-(4-Bromophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 477573-89-0P, 5-[5-(2-

(2-diethylaminoethyl)amide 477573-69-OP, 5-{5-(2-)}

Iodophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(Z)-ylidenemethyl]-2, 4dimethyl-1H-pyrcole-3-carboxylic acid (2-diethylaminoethyl)amide
477573-91-4P, 5-{5-(4-Cyanophenylmethanesulfonyl)-2-oxo-1, 2dihydroindol-3-(2)-ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic
acid (2-diethylaminoethyl)amide 477573-93-6F,
3-[[[3-[1-{4-(2-olethylaminoethylarbamoyl)-3, 5-dimethyl-1H-pyrrol-2yl]meth-(2)-ylidene]-2-oxo-2, 3-dihydro-1H-indol-5yl]sulfonyl]methyl]benzoic acid methyl ester 477573-96-9P,
2, 4-Dimethyl-5-[2-oxo-5-(3-trifluoromethoxyhenylmethanesulfonyl)-1, 2dihydroindol-3-(2)-ylidenemethyl]-1H-pyrrol-3-carboxylic acid
(2-diethylaminoethyl)amide 477573-98-IP, 5-{5-(3Cyanophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(2)-ylidenemethyl)2, 4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide
477574-00-8P, 2, 4-Dimethyl-5-(2-oxo-5-m-tolylmethanesulfonyl-1, 2dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid
(2-diethylaminoethyl)amide 477374-02-0P, 5-{5-(2-Cxo-5-m-tolylmethanesulfonyl-1, 2dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid
(2-diethylaminoethyl)amide 477374-02-0P, 5-{5-(2-(2-4-1)methyl-5-(2-0xo-5-m-tolylmethanesulfonyl-1, 2dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid
(2-diethylaminoethyl)amide 477374-02-0P, 5-{5-(2-(2-4-1)methyl-5-(2-0xo-5-m-tolylmethanesulfonyl-1, 2dihydroindol-3-(3)-ylidenemethyl-1-H-pyrrole-3-carboxylic acid
(2-diethylaminoethyl)amide 477374-02-0P, 5-{5-(2-(2-4-1)methyl-5-(2-0xo-5-m-tolylmethanesulfonyl-1, 2dihydroindol-3-(3)-ylidenemethyl-1-H-pyrrole-3-carboxylic acid

uorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl)2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide
477574-04-2P, 5-(5-(4-tert-Butylphenylmethanesulfonyl)-2-oxo-1,2dihydroindol-3-(Z)-ylidenemethyl)-Z,4-dimethyl-1H-pyrrole-3-carboxylic
acid (2-diethylaminoethyl)amide 477574-06-4P,
5-(5-(Z,6-Difluorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
(2-diethylaminoethyl)amide 477574-08-6P, 5-(5-(3Chlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl)Z,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide
477574-11-1P, 2,4-Dimethyl-5-(5-(4-mitrophenylmethanesulfonyl)-2oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid
(2-diethylaminoethyl)amide 477574-13-3P, 2,4-Dimethyl-5-(5-(3-

nitrophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl]-1H-

ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 1H-Pyrrole-3-acetamide, 5-{(z)-(5-([(2,6-dichlorophenyl)methyl)sulfonyl]-1,2-dihydro-2-xox-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-{2-(1-piperazinyl)ethyl}- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477573-60-7P, 2,4-Dimethyl-5-(2-oxo-5-phenylmethanesulfonyl-1,2-dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477573-61-8P, 5-[5-(2-Cyanophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477573-62-8P, 2,4-Dimethyl-5-[2-oxo-5-(3-tifluoromethyl)-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477573-63-0P, 5-[5-(3-Methoxyphenylmethanesulfonyl)-12-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477573-63-0P, 1ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 477573-64-P,

2,4-Dimethy1-5-[5-(2-nitrophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 477573-60-59, 2,4-Dimethyl-5-(2-oxo-5-phenylmethanesulfonyl-1,2-dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-[1,2,3]triazol-1-ylethyl)amide 477573-69-69,

2,4-Dimethyl-5-[5-(2-nitrophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-1H-pyrrole-3-carboxylic acid (2-[1,2,3]triazol-1-ylethyl)amide 477573-74-3P, 4-[[[3-[1-[4-(2-

Diethylaminoethylcarbamoyl)-3,5-dimethyl-1H-pyrrol-2-yl]meth-{2}-ylidene}2-oxo-2,3-dihydro-1H-indol-5-yl]sulfonyl]methyl]benzoic acid
478573-75-49, {4-{[[3-{[-4-{2-Diethylaminoethylcarbamoyl)-3,5-dimethyl-1H-pyrrol-2-yl]meth-{2}-ylidene}-2-oxo-2,3-dihydro-1H-indol-5-yl]sulfonyl]methyl]phenyl]acetic acid 47573-76-59,
4-{[[3-{[1-{4-{2-Diethylaminoethylcarbamoyl}}-3,5-dimethyl-1H-pyrrol-2-yl]meth-{2}-ylidene}-2-oxo-2,3-dihydro-1H-indol-5-yl]sulfonyl]methyl]a-1-ylidene

hydroxyphenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl}-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 47573-79-89,5-(5-(2-Fluorophenylmethanesulfonyl)-2-oxo-1,2-

ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-14-4P, 5-[5-(3-Bromophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-[2]-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-16-6P, 5-[5-(3,5-

Difluorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(Z)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-18-8P, 5-[5-(3,4-Difluorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl)-2,-d-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-20-2P, 5-[5-[2,5-Bis (trifluoromethyl)]-2,-d-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-22-4P, 5-[5-[3,5-Bis (trifluoromethyl)]-2,-d-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-22-4P, 5-[5-(3,5-Bis (trifluoromethyl)]-2,-d-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-24-6P,

acid (2-diethylaminoethyl)amide 477574-24-6F,

5-[5-[2-Hydroxy-5-nitrophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-26-8P, 5-[5-(2-Methoxy-5-nitrophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-28-0P, 5-[5-(2-Fluorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-30-6P, 5-[5-(3-Fluorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-32-6P, 5-[5-(4-Fluorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-35-9P, 2,4-Dimethyl-5-(2-oxo-5-(2-dihydroindol-3-(2)-ylidenemethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-35-9P, 2,4-Dimethyl-5-(2-oxo-5-(2-dihydroindol-3-(2)-ylidenemethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-35-9P, 2,4-Dimethyl-5-(2-oxo-5-(2-dihydroindol-3-(2)-ylidenemethyl)amide 477574-40-6P, 2,4-Dimethyl-3-(2-oxo-5-(2-dihydroindol-3-(2)-ylidenemethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-40-6P, 2,4-Dimethyl-3-(2-oxo-5-(2-diethylaminoethyl)amide 4775

477574-40-69, 2, 4-Dimethyl-5-(2-oxo-5
pentafluorophenylmethaneaulfonyl-1, 2-dihydroindol-3-(2)-ylidenemethyl)-1Hpyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-42-8p
, 5-[5-(2,5-Difluorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(2)ylidenemethyl)-14-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 477574-44-09, 2, 4-Dimethyl-5-(2-oxo-5(2,3,6-trifluorophenylmethanesulfonyl)-1,2-dihydroindol-3-(2)ylidenemethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide
477574-46-29, 5-(5-(2,3-Difluorophenylmethanesulfonyl)-2-oxo-1, 2dihydroindol-3-(2)-ylidenemethyl-2,4-dimethyl-1H-pyrrole-3-carboxylic
acid (2-diethylaminoethyl)amide 477574-47-3P,
5-(5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)ylidenemethyl-1-2-(4-dimethyl-1H-pyrrole-3-carboxylic acid
(2-diethylaminoethyl)amide 477574-49-5P, 5-(5-(Biphenyl-2ylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl-1-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl-1-2-oxo-1
477574-53-9P, 5-(5-(2-Fluoro-6-nitrophenylmethanesulfonyl)-2-oxo-

1, 2-dihydroindol-3-(Z)-ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 477574-53-1P, 5-[5-[2-[2-Fluorophenoxy]phenylmethanesulfonyl]-2-oxo-1, 2-dihydroindol-3-(Z)-ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid

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- ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) (2-diethylaminoethyl) amide 477574-55-39, 5-[5-(4-Chlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(2)-ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 477574-59-79, 2, 4-Dimethyl-5-(2-oxo-5-phenylmethanesulfonyl-1, 2-dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (3-diethylamino-2-hydroxypropyl) amide 477574-60-09, 2, 4-Dimethyl-5-(2-oxo-5-phenylmethanesulfonyl-1, 2-dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-(2R-tetrazol-5-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-(2R-tetrazol-5-phenylmethanesulfonyl-1, 2-dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-(2R-tetrazol-5-phenylmethanesulfonyl-1, 2-dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)-1h-pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)-1h-pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)-1h-pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)-1, 2-dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)-1h-pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)-1h-py
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid methyl(1-methylpiperidin-4-yl)amide 47757-22-77, 2-[5-[5-[2,6-]
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl)2,4-dimethyl-1H-pyrrol-3-yl)-N-[2-(3-oxopiperazin-1-yl)ethyl)acetamide
  477575-31-8P, 2-[5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo1,2-dihydroindol-3-(Z)-ylidenemethyl)-2,4-dimethyl-1H-pyrrol-3-yl)-N-[2-diethylaminoethyl)acetamide 477575-32-9P, 2-[5-(5-(2,6-
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl}-
- ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) dihydroindol-3-(Z)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-fluoroethyl)amide 477576-05-9P, 5-[5-(2,6-
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl}2,4-dimethyl-1H-pyrrole-3-carboxylic acid (3-(imidazol-1-yl)propyl)amide
  477576-06-09, 5-(5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1,2dlhydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic
  acid methylamide 477576-07-1P, 5-[5-(2,6-
- Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(Z)-ylidenemethyl)-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid amide 477576-09-3P, 5-(5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(Z)-ylidenemethyl)-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(4-acetylpiperazin-1-yl)ethyl]amide 477576-25-3P, 3-(1-(4-[[(Cyclopropyl]methylamino]methyl]-3, 5-dimethyl-1H-pyrrol-2-yl)meth-(Z)-ylidene)-5-(2,6-dichlorophenylmethanesulfonyl)-1,3-dihydroindol-2-one 477576-34-4P, 5-[5-(2,6-
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(3-acetylaminopyrrolidin-1-yl)ethyl)amide 477576-38-9P, 2-(5-[5-(2,6-
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl}-2,4-dimethyl-1H-pyrrol-3-yl)-N-[2-[4-(2-hydroxyacetyl)piperazin-1-yl]ethyl]acetamide 47757-44-6F, 2-[5-[5-[2-6]
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl)-
- 2,4-dimethyl-1H-pyrrol-3-yl]-N-[2-(2,2,2-trifluoroethylamino)ethyl]acetami de 477576-45-7P, 5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-
- de 47376-43-Py, 5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo
  1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(2,2,2-trifluoroethylamino)ethyl]amide 477576-51-58,

  5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-hydroxy)-3 (pyrrolidin-1-yl)propyl)amide 477576-52-69,

  5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (3-cyclopropylamino-2-hydroxypropyl)amide 477576-53-99,

  5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid cyclopropylamide 477576-56-09, N-[2-(3-Acetylaminopyrrolidin-1-yl)ethyl]-2-[5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrol-3-yllacetamide 477576-57-97,

  5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrol-3-yllacetamide 477576-61-79,

  2-[5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrol-3-cyllopropylamino-2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrol-3-cyllopropylamino-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrol-3-yllopropylamino-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyll-2,4-dimethyl-1H-pyrrol-3-yllopropylamino-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyll-2,4-dimethyl-1H-pyrrol-3-yllopropylamino-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyll-2,4-dimethyl-1H-pyrrol-3-yllopropylamino-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyll-2,4-dimethyl-1H-pyrrol-3-yllopropylamino-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyll-2,4-dimethyl-1H-pyrrol-3-cyllopropyl-3-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-yllopropyl-3-(3)-ylloprop
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl)-

- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-{Z}-ylidenemethyl}-2,4-dimethyl-1H-pyrrol-3-yl]-N-{Z-hydroxy-3-{1,2,3}triazol-1-ylpropyl)acetamide 477575-46-59, 5-{5-{2,6-
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-[Z]-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-methoxyethyl)amide 477375-47-6F, 5-[5-[2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-[Z]-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (3-methoxypropyl)amide 477575-48-7F, 5-[5-[2,6-
- acid (3-methoxypropyl) amide 477575-48-7p, 5-[5-(2,6Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(2)-ylidenemethyl]2,4-dimethyl-1h-pyrrole-3-carboxylic acid (2-(2-hydroxyethoxy)ethyl)amide
  477575-49-8p, 5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1, 2dihydroindol-3-(2)-ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic
  acid (2-hydroxy-1-hydroxymethyl-1-methylethyl)amide 477575-50-1p
  , 5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(2)ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid
  [2-hydroxy-1, 1-bis (hydroxymethyl) ethyl) amide 477575-57-8p,
  5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(2)ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid
  [3-(morpholin-4-yl) ethyl) amide 477575-58-8p,
  5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(2)ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid
  [3-(morpholin-4-yl) propyl) amide 477575-64-7p,
  5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(2)ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid
  [2-ethylsulfanylethyl) amide 477575-65-8p,
  5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(2)ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid
  [2-ethylsulfanylethyl) amide 477575-65-8p,
  5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(2)ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid
  [2,2,2-trifluoroethyl) amide 477575-66-3p, 5-[5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(2)ylidenemethyl-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-{Z}-ylidenemethyl]lorophenyimethanesulfonyl]-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-(pyridin-2-yl)-ethyl)amide 477575-89-6P, 5-(5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-(pyridin-3-yl)-ethyl)amide 477575-90-9P, 5-(5-(2,6-Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-(pyridin-4-yl)-ethyl) amide 477575-91-0P, 5-[5-(2,6-
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [(tetrahydrofuran-2-yl)methyl]amid 477575-92-1P, 5-[5-(Z,6-
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-{Z}-ylidenemethyl}-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (cyclopropylmethyl)amide 477576-04-89, 5-{5-{2,6-Dichlorophenylmethanesulfonyl}-2-oxo-1,2-
- ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(3-fluoropyrrolidin-1-ylethyl]amide 477577-17-69, 2-[5-[5-(2,6-
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl)-
- 2, 4-dimethyl-1H-pyrrol-3-yl]-N-[2-(3-fluoropyrrolidin-1-yl) ethyl) acetamide
  477577-26-7P, 5-[5-[2,6-Dichlorophenylmethanesulfonyl]-2-oxo-1,2dihydroindol-3-[2]-ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic
  acid [2-[4-[2-amino-2-methylproplonyl)piperazin-1-yl]ethyl]amide
  477577-31-4P, N-[2-(4-Acetylpiperazin-1-yl) ethyl]-2-[5-[5-(2,6-
- dichlorophenylmethanesulfonyl) -2-oxo-1, 2-dihydroindol-3-(Z)-ylidenemethyl)2, 4-dimethyl-1H-pyrrol-3-yl]acetamide 477577-33-69,
  5-[5-(Z, 6-Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(Z)ylidenemethyl]-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid
  [2-(4-hydroxypiperidin-1-yl)ethyl]amide 477577-35-89,
  2-[5-[5-(Z, 6-Dichlorophenylmethanesulfonyl)-2-oxo-1, 2-dihydroindol-3-(Z)ylidenemethyl]-2, 4-dimethyl-1H-pyrrol-3-yl]-N-[2-(4-hydroxypiperidin-1yl)ethyl]acetamide 477577-40-59, 5-[5-(Z, 6-
- Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid ((1-methylpiperidin-4-yl)methyl|amide 477977-42-79, 2-[5-[5-[2,6-
- yl)metnyliamide 47377-42-7F, Z-[5-[5-[2,6-]
  Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-[2)-ylidenemethyl]2,4-dimethyl-1H-pyrrol-3-yl]-N-[(1-methylpiperidin-4-yl)methyl]acetamide
  477577-54-1F, 3-[1-[4-[((Cyclopropyl)methylamino]methyl]-3,5dimethyl-1H-pyrrol-2-yl]meth(2)-ylidene]-5-[2-(2-(morpholin-4yl)ethoxy)phenylmethanesulfonyl]-1,3-dihydroindol-2-one
  477577-65-4F, 5-[5-(2,6-0ichlorophenylmethanesulfonyl)-2-oxo-1,2dihydroindol-3-[2]-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic
  acid cyclopropyl ((R)-1-pyrrolidin-2-ylmethyl) amide 477577-66-5F,
  5-[5-(2,6-0ichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-[2)ylidenemethyl]-2,4-dimethyl-1H-pyrrole3-carboxylic acid
  (cyclopropylmethyl)((R)-1-pyrolidin-2-ylmethyl)amide 477577-96-1F,
  4-(4-Fluorophenyl)-2-methyl-5-(2-oxo-5-phenylmethanesulfonyl-1,2dihydroindol-3-(2)-ylidenemethyl)-1H-pyrrole3-carboxylic acid
  (2-diethylaminoethyl)amide 477578-01-1F, 5-[5-(2,6-
- Dichlorophenylmethaneaulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
  (3-(pyrrolidin-1-yl)propyl)amide
  477578-03-39, 5-(5-(2,6-Dichlorophenylmethaneaulfonyl)-2-oxo-1,2dihydroindol-3-(Z)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic
  acid [2-(3-fluoropiperidin-1-yl)ethyl]amide 477578-07-79,
  2-(5-[5-(2,6-Dichlorophenylmethaneaulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)ylidenemethyl]-2,4-dimethyl-1H-pyrrol-3-yl]-N-[2-(3-fluoropiperidin-1yl)ethyl]acetamide
  RE: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
  (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
  (Uses)
- (drug candidate; prepn. of aralkylsulfonyl- and pyrrolylmethylidene-substituted indolinones as kinase inhibitors useful against cancers
- and
  other disorders)
  RN 477573-60-7 CAPLUS
  CN 1H-Pyrrole-3-carboxamide,
  N-[2-(diethylamino)ethyl)-5-[(2)-(1,2-dihydro-2-oxo-5-[(phenylmethyl)sulfonyl)-3H-indol-3-ylidene]methyl)-2,4-dimethyl-

ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (9CI) (CA INDEX NAME) (Continued)

Double bond geometry as shown

RN 477573-61-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-(5-[[(2-cyanophenyl)methyl]sulfonyl]-1,2dhydro-2-oxo-3H-indol-3-ylidene[methyl]-N-[2-(diethylamino)ethyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477573-62-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-(2-(diethylamino)ethyl)-5-[(2)-[1,2-dihydro-2-oxo-5-[[(3-trifluoromethyl)phenyl)methyl)sulfonyl]-3H-indol-3-ylidene|methyl]-2,4-dimethyl (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

477573-69-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-[1,2-dihydro-5-[[(2-

nitropheny1)methyl}sulfony1)-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477573-74-3 CAPLUS
CN Benzoic acid,
4-[[[(3E)-3-[(4-[[[2-(diethylamino)ethyl]amino]carbonyl]-3,5dimethyl-1H-pyrcol-2-yl]methylene)-2,3-dihydro-2-oxo-1H-indol-5yl]sulfonyl]methyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 477573-63-0 CAPLUS
COPYRIGH 2005 ACS on STN (Continued)
RN - (2-(diethylamino) ethyl) -5-((2)-(1,2-dihydro-5[(3-dihydro-5-4-dihyd

Double bond geometry as shown.

RN 477573-67-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)sthyl]-5-[(2)-(1,2-dihydro-5[(2-nitrophenyl)methyl)sulfonyl]-2-oxo-3H-indol-3-ylidene}methyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477573-68-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-[1,2-dihydro-2-oxo-5-

[(phenylmethy1)sulfony1]-3H-indol-3-ylidene]methy1)-2,4-dimethy1-N-[2-(1H-1,2,3-triazol-1-yl)ethy1]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 477573-75-4 CAPLUS
CN Benzeneacetic acid,
4-[[[(3Z)-3-[[4-[[[2-(diethylamino)ethyl]amino]carbony
1]-3,5-dimethyl-1H-pyrroi-2-yl]methylene]-2,3-dihydro-2-oxo-1H-indol-5yl]sulfonyl]methyl}- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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RN 477573-76-5 CAPLUS
CN Benzolc acid,
4-[[[(3Z)-3-[(4-[[[2-(diethylamino)ethyl]amino]carbonyl]-3,5dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-1H-indol-5yl]sulfonyl]methyl]-3-nitro- (9CI) (CA INDEX NAME)

RN 477573-78-7 CAPLUS

CN 1H-Pytrole-3-carboxamide, 5-{(Z)-[5-[{(3,5-dibromo-2-hydroxyphenyl) methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl)-N-{2-(diethylamino)ethyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477573-79-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2]-[5-[((2-fluorophenyl)methyl]sulfonyl]-1,2dinydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1H-1,2,3triazol-1-yl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 477573-82-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[[(2-chloropheny)]methyl]sulfonyl]-1,2dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-(diethylamino)ethyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477573-83-4 CAPLUS
CN Benzoic acid,
4-[[[(3Z)-3-[[4-[[[2-(diethylamino)ethyl]amino]carbonyl]-3,5dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-1H-indol-5yl]sulfonyl]methyl]-, methyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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RN 477573-84-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-{diethylamino|ethyl}-5-[(2)-[1,2-dihydro-2-oxo-5-[(4-(trifuoromethoxy)phenyl]methyl]sulfonyl]-3H-indol-3-ylidene|methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 477573-80-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl)-5-[(2)-[1,2-dihydro-4methyl-2-oxo-5-([optenylmethyl]sulfonyl]-3H-indol-3-ylidene]methyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477573-81-2 CAPLUS

RN 477573-81-2 CAPLUS

RN 477573-81-2 CAPLUS

RN 477573-81-2 CAPLUS

Fluorophenylmethyl sulfonyl]-1, 2-dihydro-4-methyl-2-oxo-3H-indol-3-ylidene]methyl]-2, 4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

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RN 477573-86-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[[(2,4-bis(trifluoromethyl)phenyl]meth
yl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-[2(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477573-88-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[[(4-bromophenyl]methyl]sulfonyl]-1,2dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-(diethylamino)ethyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

RN 477573-89-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-[(2)-[1,2-dihydro-5[[(2-iodophenyl]methyl]sulfonyl]-2-oxo-3H-indol-3-ylidene]methyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477573-91-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(Z)-[5-[(4-cyanophenyl)methyl]sulfonyl]-1,2dihydro-2-oxo-3H-indol-3-ylidene[methyl]-N-[2-(diethylamino)ethyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477573-93-6 CAPLUS

RN 4/15/3-3-0 GREAUS CN Benzoic acid, 3-[{[(3Z)-3-[[4-[[(2-(diethylamino)ethyl]amino]carbonyl]-3,5-

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 477573-98-1 CAPLUS
COPYRIGHT 2005 ACS on STN (Continued)
RN 477573-98-1 CAPLUS
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Double bond geometry as shown.

RN 477574-00-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl)-5-[(Z)-[1,2-dihydro-5[((3-methylphenyl)methyl)lsulfonyl]-2-oxo-3H-indol-3-ylidene}methyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477574-02-0 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(2)-[5-[(2,4-difluorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-1H-indol-5-yl]sulfonyl]methyl]-, methyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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RN 477573-96-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-[(Z)-[1,2-dihydro-2-oxo-5-[[[3-(trifluoromethoxy)phenyl]methyl]sulfonyl]-3H-indol-3-ylidene|methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 477574-04-2 CAPLUS 
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[{Z}-(5-[[[4-(1,1-dimethylethyl]phenyl]methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477574-06-4 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-[5-[[(2,6-difluorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-08-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-{(2)-{5-[(3)-chlorophenyl)methyl]sulfonyl]-1,2dihydro-2-oxo-3H-indol-3-ylidene]methyl}-N-[2-(diethylamino)ethyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477574-11-1 CAPLUS

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L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-[(2]-[1,2-dihydro-5[[(4-nitrophenyl]methyl]sulfonyl]-2-oxo-3H-indol-3-ylidene]methyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-13-3 CAPLUS
CN 1H-Pytrole-3-carboxamide,
N-[2-(dichylamino)ethyl]-5-[(2)-[1,2-dihydro-5[((3-nitrophenyl)methyl)sulfonyl]-2-oxo-3H-indol-3-ylidene)methyl]-2,4dimethyl- (9c1) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-14-4 CAPLUS
CN 1H-Pyrcole-3-carboxamide,
5-[(2)-[5-[(3)-bromophenyl)methyl]sulfonyl]-1,2dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continue A77574-20-2 CAPLUS COPYRIGHT 2005 ACS on STN (Continue CONTINUE A77574-20-2 CAPLUS COPYRIGHT 2005 ACS ON STN (Continue CONTINUE A77574-20-2 CAPLUS COPYRIGHT 2005 ACS ON STN (Continue CONTINUE A77574-2005)

5-{(2)-{(5)-{(12)-{(12)-{(5)-{(12)

Double bond geometry as shown.

477574-22-4 CAPLUS

1H-Pyrrole-3-carboxamide,

(2) -[5-[[3,5-bls(trifluoromethyl)phenyl]meth
y]sulfonyl]-1,2-dhydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-24-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-{2-(diethylamino)ethyl)-5-{(Z)-[1,2-dihydro-5[{(2-hydroxy-5-nitrophenyl)methyl)aulfonyl]-2-oxo-3H-indol-3ylidene|methyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

477574-16-6 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[{Z}-[5-[{(3,5-difluorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477574-18-8 CAPLUS 1H-Pyrrole-3-carboxamide, N-{2-(diethylamino)ethyl]-5-{Z}-{5-{[(3,4-difluorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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477574-26-8 CAPLUS
1H-Pyrrole-3-carboxamide,
-(diethylamino)ethyl]-5-((2)-(1,2-dihydro-5[(2-methoxy-5-ntrophenyl)methyl]sulfonyl]-2-oxo-3H-indol-3ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477574-28-0 CAPLUS 1H-Pyrrole-3-carboxamide,  $N-\{2-\{diethylamino\}ethyl\}-5-\{\{Z\}-\{5-\{\{(2-1)^2\}-1\}\}-1\}\}$ 

fluorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl}2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477574-30-4 CAPLUS 1H-Pyrrole-3-carboxamide, N-{2-(diethylamino)ethyl]-5-[(Z)-[5-[((3-

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fluorophenyl)methyl|sulfonyl|-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl|
2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477574-32-6 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-{diethylamino}ethyl]-5-[{Z}-[5-[[{4-

fluorophenyl)methyl}sulfonyl}-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl}2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-35-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-[(2)-[1,2-dihydro-2oxo-5-[[(2-(trifuoromethyl)phenyl]methyl]sulfonyl]-3H-indol-3ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

477574-42-8 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[{2}-{5-{{(2,5-difluorohenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene}methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-44-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-[{Z}-[1,2-dihydro-2-

oxo-5-[{(2,3,6-trifluorophenyl)methyl}sulfonyl}-3H-indol-3-ylidene}methyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477574-46-2 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-[5-[((2,3-difluorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 477574-37-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl)-5-[(Z)-[1,2-dihydro-2-oxo-5-[[4-(trifluoromethyl)phenyl]methyl]=ulfonyl]-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-40-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-{2-(diethylamino)ethyl}-5-[{Z}-[1,2-dihydro-2-

oxo-5-[[(pentafluorophenyl)methyl]sulfonyl]-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

RN 477574-47-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(Z)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]-

1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-{2-(diethylamino)ethyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-49-5 CAPLUS CN 1H-Pyrrole-3-carboxamide, 5-[(Z)-[5-[([1,1'-bipheny1]-2-ylmethy1)sulfony1]-

1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-51-9 CAPLUS
CN 1H-Fyrrole-3-carboxamide,
N-[2-(diethylamino) ethyl]-5-[(2)-[5-[(2-fluoro-6-nitrophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-

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ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME) (Continued)

Double bond geometry as shown.

477574-53-1 CAPLUS  $\frac{1}{1} - \frac{1}{2} - \frac{1}$ 

Double bond geometry as shown.

RN 477574-55-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[[(4-chloropheny]]methyl]sulfonyl]-1,2dhydro-2-oxo-3h-indol-3-ylidene|methyl]-N-[2-(diethylamino)ethyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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477574-65-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{Z}-[1,2-dihydro-2-oxo-5-[(phenylmethyl)sulfonyl]-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-66-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-[bis[1-methylethyl] amino]ethyl]-5-[[Z]-[1,2-dinyloro-2-oxo-5-[[phenylmethyl] sulfonyl]-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-67-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2]-[5-[([2-fluorophenyl)methyl]sulfonyl]-1,2dhydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1pyrrolidinyl)ethyl)- (9CI) (CA INDEX NAME)

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 477574-59-7 CAPLUS
CO 1H-Pyrrole-3-carboxamid+,
N-[3-(diethylamino)-2-hydroxypropyl]-5-{(Z)-{1,2-diethylamino}-2-hydroxypropyl]-5-{(Z)-{1,2-diethylamino}-2-hydroxypropyl]-3-diethylaulfonyl]-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

477574-60-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-((Z)-(1,2-dihydro-2-oxo-5-

{ (phenylmethyl) sulfonyl } -3H-indol-3-ylidene]methyl}-2,4-dimethyl-N-[2-{1H-tetrazol-5-yl}ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-64-4 CAPLUS

N H-Pyrrole-3-carboxamide,
N-[2-diethylamino|ethyl]-5-[(z)-[1,2-dihydro-2-oxo-5-[(phenylmethyl)sulfonyl]-3H-indol-3-ylidene|methyl]-4-methyl- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) Double bond geometry as shown.

477574-68-8 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-[5-[[(2-

fluorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl}4-methyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477574-70-2 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-{bis(1-methylethyl)amino}ethyl]-5-{(2)-[5-[(2-fluorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene}methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 477574-74-6 CAPLUS
COPYRIGH 2005 ACS on STN (Continued)
CONTINUE CONTIN

Double bond geometry as shown.

RN 477574-77-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(3)-(5-[(3-chlorophenyl)methyl)sulfonyl]-1,2dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(1pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477574-78-0 CAPLUS
1H-Pytrole-3-carboxamide, N-[2-[bis(1-methylethyl)amino]ethyl]-5-{(Z)~[5-[([3-chlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

477574-90-6 CAPLUS
1H-Pyrrole-3-carboxamide,
(2) -[5-[[(2,6-dichlorophenyl]methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(3-oxo-1-piperazinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477575-11-4 CAPLUS
CN HH-Pytrole-3-carboxamide,
5-(2)-15-[(2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl)-N-(2-hydroxyethyl)-2,4dimethyl- (SCI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

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RN 477574-88-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477574-89-3 CAPLUS
CN 1H-Pytrole-3-carboxamide,
5-[(2)-15-[(2,6-61-cho)rophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl- (SCI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 477575-16-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(Z)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-

1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N,2,4-trimethyl-N-(1-methyl-4-piperidinyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477575-22-7 CAPLUS HI-Fyrrole-3-acetamide, 5-[(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(3-oxo-1-piperazinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

 $\begin{array}{lll} 477575-31-9 & CAPLUS & \\ 1H-Pyrrole-3-acetamide, & 5-\{\{Z\}-\{5-\{\{\{2,6-dichlorophenyl\}methyl\}sulfonyl\}-1\}-1\}-1 & \\ 1H-Pyrrole-3-acetamide, & 5-\{\{Z\}-\{\{1,6-dichlorophenyl\}methyl\}sulfonyl\}-1 & \\ 1H-Pyrrole-3-acetamide, & 5-\{\{Z\}-\{\{1,6-dichlorophenyl\}methyl\}-1 & \\ 1H-Pyrrole-3-acetamide, & 5-\{\{1,6-dichlorophenyl\}methyl]-1 & \\ 1H-Pyrrole-3-acetamide, & 5-\{\{1,6-dichlorophenyl\}methyllorophenyllor$ 1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

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477575-32-9 CAPLUS  $lH-Pyrrole-3-acetamide, \ 5-[(Z)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-1 + (2,6-dichlorophenyl)methyl [sulfonyl] - (3,6-dichlorophenyl)methyl [sulfonyl] - (4,6-dichlorophenyl)methyl [sulfonyl] - (4,6-dichlorophenyl] - (4,6-dichlorophe$ 

-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N,2,4-trimethyl-N-(1-methyl-4-piperidinyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477575-34-1 CAPLUS

IH-Pyrrole-3-acetamide, 5-[(Z)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-y)rrolidinyl)ethyl]- (GA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 477575-46-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-{[2]-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylldene]methyl]-N-(2-methoxyethyl)-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477575-47-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[((2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-(3-methoxypropyl)-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477575-48-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

477575-37-4 CAPLUS
1H-Pyrrole-3-acetamide, 5-[(2)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-hydroxy-3-[4morpholinyl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477575-38-5 CAPLUS
1H-Pyrrole-3-acetamide, 5-[(2)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-[2-(2-hydroxyethoxy)ethyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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RN 477575-49-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[(2,6-dichloropheny)]methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-hydroxy-1(hydroxymethyl)-1-methylethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477515-50-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-hydroxy-1,1-bis(hydroxymethyl)ethyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

RN 477575-57-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-(22)-(5-[(2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-yiidene|methyl]-2,4-dimethyl-N-{2-(4-morpholinyl)ethyl}- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477575-58-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-((2)-(5-[((2,6-dichlorophenyl)methyl]sulfonyl)1,2-dinydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[3-(4-morpholinyl)propyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl|-2,4-dimethyl-N-[2-(2-pyridinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477575-89-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2]-(5-[((2,6-dichlorophenyl)methyl)sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(3-pyridinyl)ethyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477575-90-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(4-pyridinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 477575-64-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-(5-[(2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-{ethylthio}ethyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477575-65-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[(2,6-dichlorophenyl)methyl)sulfonyl)1,2-dihydro-2-oxo-3H-indol-3-ylidenejmethyl)-2,4-dimethyl-N-(2,2,2-trifluoroethyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477575-86-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(Z)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 477575-91-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[([2,6-dichlorophenyl)methyl]sulfonyl]-

1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[(tetrahydro-2-furanyl)methyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

AN 477575-92-1 CAPLUS

AN H-Pyrrole-3-carboxamide, N-(cyclopropylmethyl)-5-{(Z)-[5-{((2,6-dichlorophenyl))methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 477576-04-8 CAPLUS
COPYRIGHT 2005 ACS on STN (Continued)
RN 477576-04-8 CAPLUS
(1) -[5-[(12,6-dichlorophenyl)methyl]sulfonyl)[2,2-dihydro-2-ox-3H-indol-3-ylidene]methyl]-N-(2-fluoroethyl)-2,4dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477576-05-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[(2,6-dichlorophenyl]methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[3-(1H-imidazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477576-06-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-(12)-5-(12,6-6-(10,6-6-(10-cophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N,2,4-trimethyl- (9CI) (CA
INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

477576-25-3 CAPLUS
2H-Indol-2-one, 3-[(4-[(cyclopropylmethylamino)methyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-1,3-dihydro-, (3Z)- [9CI) (CA INDEX NAME)

Double bond geometry as shown.

477576-34-4 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-[3-(acetylamino)-1-pyrrolidinyl]ethyl]-5-

[(2)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 477576-07-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-{(Z)-(5-{((2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477576-09-3 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(4-acetyl-1-piperazinyl)ethyl]-5-[(2)-[5-([2,6-dichlorophenyl)methyl]sulfonyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

477576-38-8 CAPLUS lH-Pyrrole-3-acetamide,  $5-[(2)-[5-[\{(2,6-dichloropheny1)methy1\}sulfony1)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-[4-(hydroxyacety1)-1-piperaziny1]ethyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)$ 

Double bond geometry as shown.

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477576-44-6 CAPLUS  $\begin{array}{lll} & & & & \\ & 1 \\ & 1 \\ & 1 \\ & 1 \\ & 1 \\ & 2 \\ & 1 \\ & 1 \\ & 2 \\ & 2 \\ & 1 \\ & 2$ 

Double bond geometry as shown.

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RN 477576-45-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2]-(5-[(2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylldene|methyl]-2,4-dimethyl-N-[2-[(2,2,2-trifluoroethyl)amino]ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477576-51-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(Z)-[5-[(Z)-6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-hydroxy-3-(1-pyrrolidinyl)propyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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RN 477576-57-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[(2),6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidenelmethyl]-N-[2-[4-(hydroxyacetyl)-1-piperazinyl]ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 477576-52-6 CAPLUS
COPYRIGHT 2005 ACS on STN (Continued)
N-[3-(cyclopropylamino)-2-hydroxypropyl]-5-[(z)[5-[(z,6-dichlorophenyl)methyl]aulfonyl]-1,2-dihydro-2-oxo-3H-indol-3ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown

N 477576-55-9 CAPLUS
N 1H-Pyrrole-3-carboxamide, N-cyclopropyl-5-[(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

Double bond geometry as shown.

RN 477576-56-0 CAPLUS
CN 1H-Pyrrole-3-acetamide,
N-[2-[3-[acetylamino]-1-pyrrolidinyl]ethyl]-5-[(z)[5-[([2,6-dichlorophenyl]methyl]aulfonyl]-1,2-dihydro-2-oxo-3H-indol-3ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued

RN 477576-61-7 CAPLUS
CN 1H-Pyrrole-3-acetamide, 5-[(2)-[5-[(2,6-dichloropheny1)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-(2-hydroxy-3-(1pyrrolidinyl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477576-62-8 CAPLUS
CN 1H-Pytrole-3-acetamide,
N-[3-(cyclopropylamino)-2-hydroxypropyl]-5-[(z)-[5[[(2,6-dichlorophenyl)mathyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3ylidene|methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477576-95-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-(4-fluoro-1-piperidinyl)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477576-98-0 CAPLUS
1H-Pyrrole-3-acetamide, 5-[(2)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-(4-fluoro-1-piperidinyl)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

# Double bond geometry as shown.

RN 477577-16-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2]-[5-[((2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-(3-fluoro-1pyrrolidinyl)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

## Double bond geometry as shown.

## L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

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NH2

477577-31-4 CAPLUS HH-Pyrrole-3-acetamide, N-[2-(4-acetyl-1-piperaziny1)ethyl]-5-[(2)-[5-[(2,6-dich)eropheny1]methyl]sulfonyl]-1, 2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2, 4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477577-33-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(Z)-[5-[[(2,6-dichlorophenyl)methyl]aulfonyl]-

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

Double bond geometry as shown.

477577-26-7 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-[4-(2-amino-2-methyl-1-oxopropyl)-1-piperazinyllethyl]-5-[(2)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- [9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-{4-hydroxy-1-piperidinyl}ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477577-35-8 CAPLUS
1H-Pyrrole-3-acetamide, 5-[(Z)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-(4-hydroxy-1-piperidinyl)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

PAGE 1-B

RN 477577-40-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(Z)-[5-[((2,6-dichlorophenyl)methyl]sulfonyl]-

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ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[{l-methyl-4-piperidinyl}methyl}- (9CI) (CA INDEX NAME)

477577-42-7 CAPLUS 1H-Pyrrole-3-acetamide, 5-[(2)-[5-[((2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidenejmethyl]-2,4-dimethyl-N-[((1-methyl-4-piperidinyl)methyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

 $\begin{array}{lll} 477577-54-1 & CAPLUS \\ 2H-Indol-2-one, & 3-[\{4-\{(cyclopropylmethylamino)methyl\}-3,5-dimethyl-1H-pyrrol-2-yllmethylene]-1,3-dihydro-5-\{\{[2-\{2-\{4-morpholinyl)ethoxy]phenyl]methyl]sulfonyl]-, & (32)- & (9CI) & (CA INDEX NAME) & (32)- & (9CI) & (CA INDEX NAME) & (32)- & (9CI) & (CA INDEX NAME) & (32)- & (9CI) &$ 

Double bond geometry as shown.

ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

RN 477577-96-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-[(Z)-[1,2-dihydro-2-oxo-5-[(phenylmethyl)sulfonyl]-3H-indol-3-ylidene)methyl]-4-(4-fluorophenyl)-2-methyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477578-01-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-(2)-(5-[((2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[3-(1-pyrrolidinyl)propyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 477578-03-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-(5-[(2,6-dichlorophenyl)methyl]sulfonyl)1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(3-fluoro-1piperidinyl)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

477577-65-4 CAPLUS

1H-Pyrrole-3-carboxamide, N-cyclopropyl-5-[(Z)-{5-[(Z,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[(ZR)-2-pyrrolidinylmethyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

477577-66-5 CAPLUS
1H-Pyrrole-3-carboxamide, N-(cyclopropylmethyl)-5-[(Z)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[(2R)-2-pyrrolidinylmethyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

Double bond geometry as shown.

477578-07-7 CAPLUS
1H-Pyrrole-3-acetamide, 5-[(2)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(3-fluoro-1-piperidinyl)ethyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

477576-37-7P, 4-[2-[[2-[5-[5-(2,6-Dichlorophenylmethanesulfonyl]-2-oxo-1,2-dihydroindol-3-(2]-ylidenemethyl]-2,4-dimethyl-1H-pyrrol-3-yl]acetyl]amino]ethyl]piperarine-1-carboxylic acid tert-butyl ester 477576-39-9P, Acetic acid 2-[4-[2-[[2-[5-[5-(2,6-

dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl]-2,4-dimethyl-1H-pyrrol-3-yl]acetyl]amino]ethyl]piperazin-1-yl]-2-oxoethyl ester 477576-58-2P, 4-[2-[[5-[5-(2,6-

Dichlorophenylmethanesulfonyl)-2-oxo-1,2-dihydroindol-3-(2)-ylidenemethyl]-2,4-dimethyl-1H-pyrrol-3-ylcarbonyl]amino|ethyl]piperazine-1-carboxylic acid tert-butyl ester 477576-59-3P, 5-[5-(2,6-

Dichlorophenylmethanesulfonyl}-2-oxo-1,2-dihydroindol-3-(Z)-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-(piperazin-1-yl)ethyl)amide 477576-60-6P, Acetic acid 2-[4-[2-[(5-[5-(2,6-

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 $\label{local-equation} \mbox{dichlorophenylmethanesulfonyl} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl\} - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl] - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl] - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl] - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl] - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyl] - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyll - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyll - 2 - oxo - 1, 2 - dihydroindol - 3 - \{Z\} - ylidenemethyll - 2 - oxo - 1, 2 - dihydroindol - 3 - (Z) - ylidenemethyll - 2 - oxo - 1, 2 - dihydroindol - 3 - (Z) - ylidenemethyll - 2 - oxo - 1, 2 - dihydroindol - 3 - (Z) - ylidenemethyll - 2 - oxo - 1, 2 - dihydroindol - 3 - (Z) - ylidenemethyll - 2 - oxo - 1, 2 - dihydroindol - 3 - (Z) - ylidenemethyll - 2 - oxo - 1, 2 - dihydroindol - 3 - (Z) - ylidenemethyll - 2 - oxo - 1, 2 - dihydroindol - 3 - (Z) - ylidenemethyll - 3 - (Z) - (Z$ 

2,4-dimethyl-1H-pyrrol-3-ylcarbonyl]amino]ethyl]piperazin-1-yl}-2-oxoethyl ester 477577-27-89, [2-[4-[2-[[5-[5-[42,6-

Dichlorophenylmethanesulfonyl) -2-oxo-1, 2-dihydroindol-3-(Z)-ylidenemethyl)2, 4-dimethyl-1H-pyrrol-3-ylcarbonyl amino) ethyl piperaxin-1-yl]-1, 1dimethyl-2-oxoethyl carbamic acid tert-butyl ester
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
(Reactant or reagent)
(prepn. of aralkylsulfonyl- and pyrrolylmethylidene-substituted
indolinones as kinase inhibitors useful against cancers and other
disorders)
RN 477576-37-7 CAPLUS
CN 1-Piperazinecarboxylic acid, 4-[2-[[[5-[(Z)-[5-[(Z)-6dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3ylidene[methyl]-2,4-dimethyl-1H-pyrcol-3-yllace(yllamino)ethyl]-,
1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

PAGE 1-E

\_OBu−t

RN 477576-39-9 CAPLUS
CN 1H-Pyrrole-3-acetamide,
N-[2-[4-[(acetyloxy)acetyl]-1-piperazinyl]ethyl]-5-

[(2)-[5-[[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

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-oBu-t

RN 477576-59-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(1-piperazinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

477576-60-6 CAPLUS
1H-Pyrrole-3-carboxamide,
-[4-[(acetyloxy)acetyl]-1-piperazinyl]ethyl]5-[(2)-[5-[(12,6-dichlorophenyl)methyl]sulfonyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene|methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) PAGE 1-A

PAGE 1-B

477576-58-2 CAPLUS
1-Piperazinecarboxylic acid, 4-[2-[[[5-[(Z)-[5-[{(Z,6-dichlorophenyl)methyl]=0,4-dinethyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidenelmethyl]-2,4-dimethyl1-H-pyrcol-3-yl]carbonyl]amino]ethyl]-,1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 53 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

PAGE 1-B

-OAC

477577-27-8 CAPLUS

Double bond geometry as shown.

PAGE 1-B

ANSWER 54 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

RN 356069-35-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 54 OF 65 CAPLUS COPYRIGHT 2005 ACS ON STN ACCESSION NUMBER: 2002:805222 CAPLUS COPUMENT NUMBER: 139:270353

ACCESSION NUMBER: 202:805222 CAPLUS

DOCUMENT NUMBER: 139:270353

INITIE: Inhibition of constitutively active forms of mutant kit by multitargeted indolinone tyrosine kinase inhibitors. [Erratum to document cited in CAI38:147266]

AUTHOR(S): Liao, Albert T.: Chien, May B.: Shenoy, Narmads; Mendel, Dirk B.: McMahon, Gerald: Cherrington, Julie M.: London, Cheryl A.

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SOURCE: Bloody ISSN: 0006-4971

PUBLISHER: American Society of Hematology
DOCUMENT TYPE: Journal
LANGUAGE: Journal
LAN

Double bond geometry as shown.

NAME)

Double bond geometry as shown.

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

| PATENT NO.    |       |     |      |           |     |          |      |                 | APPLICATION NO. |      |      |      |     |          |      |      |     |
|---------------|-------|-----|------|-----------|-----|----------|------|-----------------|-----------------|------|------|------|-----|----------|------|------|-----|
|               |       |     |      |           |     |          |      |                 |                 |      |      |      |     |          |      |      |     |
| WO 2002081466 |       |     |      | A1 200210 |     |          | 1017 | WO 2002-US11001 |                 |      |      |      |     | 20020409 |      |      |     |
|               | W:    | ΑE, | AG,  | AL,       | AM, | ΑT,      | ΑU,  | ΑZ,             | BA,             | ₿₿,  | BG,  | BR,  | BY, | BZ,      | CA,  | CH,  | CN, |
|               |       | co, | CR,  | CU,       | CZ, | DΕ,      | DK,  | DM,             | DZ,             | EC,  | EE,  | ES,  | FI, | GB,      | GD,  | GE,  | GH, |
|               |       | GM, | HR,  | ΗU,       | ID, | IL,      | IN,  | IS,             | JP,             | ΚE,  | KG,  | KP,  | KR, | KZ,      | LC,  | LK,  | LR, |
|               |       | LS, | LT,  | LU,       | LV, | ΜA,      | MD,  | MG,             | MK,             | MN,  | MW,  | MX,  | MZ, | NO,      | NZ,  | OM,  | PH, |
|               |       | PL, | PT,  | RO,       | RU, | SD,      | SE,  | SG,             | SI,             | SK,  | SL,  | TJ,  | TM, | TN,      | TR,  | TT,  | TZ, |
|               |       | UΑ, | UG,  | US,       | UΖ, | VN,      | YU,  | ZA,             | ZM,             | ZW,  | AM,  | ΑZ,  | BY, | KG,      | KZ,  | MD,  | RU, |
|               |       | ŦJ, | TM   |           |     |          |      |                 |                 |      |      |      |     |          |      |      |     |
|               | RW:   | GH, | GM,  | KE,       | LS, | MW,      | MZ,  | SD,             | SL,             | SZ,  | ΤZ,  | UG,  | ZM, | ZW,      | AT,  | BE,  | CH, |
|               |       |     |      |           |     |          | FR,  |                 |                 |      |      |      |     |          |      |      |     |
|               |       |     |      | CF,       | CG, |          | CM,  |                 |                 |      |      |      |     |          |      |      |     |
| US 2003100555 |       |     |      |           |     |          |      |                 | US 2002-118321  |      |      |      |     | 20020409 |      |      |     |
| US 6797725    |       |     |      |           |     | 2004     |      |                 |                 |      |      |      |     |          |      |      |     |
| US 2004186161 |       |     |      | A1        |     | 20040923 |      |                 |                 |      |      |      |     |          |      |      |     |
| PRIORIT       | Y APP | LN. | INFO | .:        |     |          |      |                 |                 | US 2 | 001- | 2826 | 30P |          | P 2  | 0010 | 409 |
|               |       |     |      |           |     |          |      |                 | ,               | US 2 | 002- | 1183 | 21  |          | A3 2 | 0020 | 409 |

OTHER SOURCE(S): MARPAT 137:294870

The present invention relates to pyrrole substituted 2-indolinone compds. (shown as I; e.g.  $3-\{1-(3,5-\text{dimethyl-1H-pyrrol-2-yl)meth-(2)-ylidene}-2-\cos 2,3-\text{dihydroindole-1-carbonyl chloride})$  and their pharmaceutically

Page 78 SAEED ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) acceptable salts which modulate the activity of protein kinases and therefore are expected to be useful in the prevention and treatment of protein kinase related cellular disorders such as cancer (no data). In

R1 and R2 are independently H, halo, alkyl, alkylthio, nitro, trihalomethyl, hydroxy, hydroxyalkyl, alkoxy, cyano, aryl, heteroaryl, -c(0)R7 (R7 is alkyl, amino, hydroxy, alkoxy, aryl, heteroaryl, aryloxy, heteroaryloxy, heterocycle, and aminoalkylamino), -NR8R9, -RR8C(0)R9, -SO2R8, and -S(0)2NR8R9 (R8 and R9 are independently H, alkyl, aryl and heteroaryl, or R8 and R9 together with the N to which they are attached form a satd. heterocycloamino). R3 is H, alkyl, hydroxyalkyl,

aminoalkyl,

--(O)R7, aryl, and heteroaryl; R4 is H, alkyl, --C(O)R7 aryl, and
heteroaryl. R5 is H and --COR10 where R10 is alkyl, alkoxy, hydroxy,

heteroaryl. R5 is H and -COR10 where R10 is alkyl, alkoxy, hydroxy, arylox, heteroaryl, heterocycle, alkylamino, dialkylamino, or -NRIIR12 where R11 is H or alkyl, and R12 is aminoalkyl, hydroxyalkyl, carboxyalkyl, alkoxycarbonylalkyl, hydroxyalkyl, or heterocyclylalkyl wherein the alkyl chain in aminoalkyl, heteroaralkyl, heteroaralkyl, or heterocyclylalkyl is optionally substituted with one or two hydroxy group(s); or R4 and R5 together form - (CR2)4- or -(CR2)mCO(CR2)n- wherein n is O to 3, provided that n+m is 3. R6 is: (c) -OR13 wherein R13 is alkyl, trifluoromethyl, carboxyalkyl, aminoalkyl, phosphonooxyalkyl, sulfooxyalkyl, hydroxyalkyl, alkoxyalkyl, aryl, heterocaryl, heteroaralkyl, heterocyclyl, monoaccharides and heterocyclylalkyl, sulfooxyalkyl, that heteroaralkyl, carboxyalkyl, sulfooxyalkyl, hydroxyalkyl, heterocyclylalkyl, hydroxyalkyl, or alkoxyalkyl is optionally substituted with one or two hydroxy group(s) and further wherein one or two C atoms in said alkyl chain are optionally replaced by O, -NR14- (R14 is H or alkyl), -S-, or -SO2-; or. (d) -NR15R16 where are R15 and R16 are independently H, alkyl,

carboxyalkyl, alkoxyalkyl, aminoalkyl, phosphonooxyalkyl, sulfooxyalkyl, hydroxyalkyl, aryl, heteroaryl, heteroaralkyl, and heterocyclylalkyl; wherein the alkyl chain in carboxyalkyl, aminoalkyl, phosphonooxyalkyl, heteroaralkyl, heterocyclylalkyl, hydroxyalkyl, or alkoxyalkyl is optionally substituted with one or two hydroxy group(s) and further wherein one or two C atoms in the alkyl chain are optionally replaced by O, -NR17- (R17 is H or alkyl), -5-, or -502-; or R15 and R16 together with

o, -MRI'- (RI7 is H of airyi), -s-, or -soc-; of KIS and RIO Cogener

the N atom to which they are attached form satd. or unsatd.
heterocycloamino;. Although the methods of prepn. are not claimed, >80
example prepns. are included, both of I and the unprotected version of I
in which the C(O)R6 group has been replaced by H.

IT 356086-94-5p, 5-(5-Fluoro-2-oxo-1,2-dihydroindol-(32)ylidenmethyll-2,4-dimethyl-1H-pyrole-3-carboxylic acid
(2-(pyrrolidin-1-yl)ethyllamide
RL: RCT (Reactant): SPN (Synthetic preparation); PREP (Preparation); RACT
(Reactant or reagent)
(intermediate; preparation of prodrugs of
(pyrrolylmethylidenelpindolinones
and activity as modulators of protein kinases)
RN 356088-94-5 CAPJUS
CN 1H-Pyrrole-3-carboxamide, 5-(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 342641-52-5P

Paromo-2-oxo-1, 2-dihydroindol-3-ylidenemethyl)-2-isopropyl-4-phenyl1H-pyrrole-3-carboxylic acid [3-(4-methylpiperaxin-1-yl)propyl) amide
342641-54-7P, 5-(5-Bromo-2-oxo-1, 2-dihydroindol-3-ylidenemethyl)-2methyl-4-phenyl-1H-pyrrole-3-carboxylic acid (2-(pyrolidin-1yl)ethyl)amide 342641-55-6P, 5-[6-(2-Methoxyphenyl)-2-oxo-1, 2dihydroindol-3-ylidenemethyl]-2-methyl-4-phenyl-1H-pyrrole-3-carboxylic
acid (2-(pyrrolidin-1-yl)ethyl)amide 342641-56-9P,
5-(5-Bromo-2-oxo-1, 2-dihydroindol-3-ylidenemethyl)-2-methyl-4-phenyl-1Hpyrrole-3-carboxylic acid (2-dimethylaminoethyl)amide 342641-57-0P

acid (2-Bromo-2-oxo-1, 2-dihydroindol-3-ylidenemethyl)-2-methyl-4-phenyl-1H-pyrrole-3-carboxylic acid (2-dimethylaminoethyl) amide 34264-57-09

5-[6-(2-Methoxyphenyl)-2-oxo-1, 2-dihydroindol-3-ylidenemethyl)-2-methyl-4-phenyl-1H-pyrrole-3-carboxylic acid (2-dimethylaminoethyl) amide 34264-59-2P, 5-(5-Bromo-2-oxo-1, 2-dihydroindol-3-ylidenemethyl)-2-methyl-4-phenyl-1H-pyrrole-3-carboxylic acid (3-diethylaminoethyl) amide 34264-60-5P, 5-(5-Bromo-2-oxo-1, 2-dihydroindol-3-ylidenemethyl)-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid (2-dimethylaminoethyl) amide 34264-60-5P, 5-(5-Bromo-2-oxo-1, 2-dihydroindol-3-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-dimethylaminoethyl) amide 34264-61-6P, 2, 4-Dimethyl-5-(2-oxo-6-phenyl-1, 2-dihydroindol-3-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (2-dimethylaminoethyl) amide 34264-63-6P, 5-(5-Bromo-2-oxo-1, 2-dihydroindol-3-ylidenemethyl)-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid (2-dimethylaminoethyl) amide 342641-63-8P, 5-(5-Bromo-2-oxo-1, 2-dihydroindol-3-ylidenemethyl)-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)ethyl)amide 342641-65-9P, 5-(5-Bromo-2-oxo-1, 2-dihydroindol-3-ylidenemethyl)-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid (1-dimethyl-1H-pyrrole-3-carboxylic acid (2-dimethyl-1H-pyrrole-3-carboxylic acid (2-dimethyl-1)-2, 4-dimethyl-1H-pyrrole-3-carboxylic acid (2-dimethyl-1)-2, 4-dimethyl-1-3-ylidenemethyl)-2, 4-dimethyl-1-3-ylidenemethyl)-2, 4-dimethyl-1-3-ylidenemethyl-2-2, 4-Dimethyl-5-(2-oxo-5-phenyl-1, 2-dihydroindol-3-ylidenemethyl)-1-1-pyrrole-3-carboxylic acid (2-dimethylaminoethyl) amide 342641-66-3P, 2, 4-Dimethyl-5-(2-oxo-5-phenyl-1, 2-dihydroindol-3-ylidenemethyl)-1-1-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 342641-69-4P, 2, 4-Dimethyl-5-(2-oxo-5-phenyl-1, 2-dihydroindol-3-ylidenemethyl)-1-1-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 342641-70-7P, 2, 4-Dimethyl-5-(2-oxo-6-phenyl-1, 2-dihydroindol-3-ylidenemethyl)-1-1-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 342641-70-7P, 2, 4-Dimethyl-5-(2-oxo-6-

2,4-Dimethyl-5-(2-oxo-6-pyridin-3-yl-1,2-dihydroindol-3-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (3-dimethylaminopropyl)amide 34264-78-5p, 2,4-Dimethyl-5-(2-oxo-5-phenyl-1,2-dihydroindol-3-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (3-dimethylaminopropyl)amide

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA NAME)

Double bond geometry as shown.

S57795-19-4P, 5-(5-Fluoro-2-oxo-1,2-dihydroindol-(3Z)ylidenemethyl)-2,4-dimethyl-lH-pyrrole-3-carboxylic acid
(2-diethylaminoethyl)amide
RE: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic
preparation); THU (Therapeutic use); BIOL (Biological study); PREP
(Preparation); RACT (Reactant or reagent); USES (Uses)
(protein kinase modulator; preparation of prodrugs of
(pyrrolylmethylidene)indolinones and activity as modulators of protein
kinases)
557795-19-4 CAPLUS
H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX
NAME)

Double bond geometry as shown.

$$\begin{array}{c|c} & & & \\ & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ &$$

342641-49-09, 5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2-isopropyl-4-phenyl-1H-pyrrole-3-carboxylic acid (3-diethylaminopropyl) amide 342641-50-39, 5-(5-Bromo-2-oxo-1,2-

5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2-isopropyl-4-phenyl-1H-

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 342641-79-69, 2,4-Dimethyl-5-(2-oxo-5-phenyl-1,2-dihydroindol-3-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (3-diethylaminopropyl) amide 342641-80-99, 2,4-Dimethyl-5-(2-oxo-6-phenyl-1,2-dihydroindol-3-ylidenemethyl)-1H-pyrrole-3-carboxylic acid (3-diethylaminopropyl) amide 342641-81-09, 3-[4-(3-Diethylaminopropylcarbamoyl)-3,5-dimethyl-1H-pyrrole-2-oxo-2,3-dihydro-1H-indole-4-carboxylic acid (3-chloro-4-methoxyphenyl) amide 342641-82-19,

(3-cloro-4-metnoxypneny); amide 34284-92-19,

-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole3-carboxylic acid (3-diethylaminopropyl)amide 342641-93-29,

5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-12,4-diisopropyl-1Hpyrrole-3-carboxylic acid (2-diethylaminopropyl)amide 342641-04-39

,5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-diisopropyl-1Hpyrrole-3-carboxylic acid (3-diethylaminopropyl)amide 342641-05-49

,5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-diisopropyl-1Hpyrrole-3-carboxylic acid (3-(pyrrolidin-1-yl)propyl)amide
342641-07-69, 5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)2,4-dimethyl-1H-pyrrole-3-carboxylic acid (pyrtidin-4-ylmethyl)amide
342641-08-79, 5-(6-(4-Butylphenyl)-2-oxo-1,2-dihydroindol-3ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
(2-(pyrrolidin-1-yl)ethyl)amide 342641-09-09,

(z-(pyrrolidin-1-yl)ethyl)amide 342641-89-8F,

5-[6-(5-Isopropyl-2-methoxyphenyl)-2-oxo-1,2-dihydroindol-3-ylidenemethyl]2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)ethyl)amide
342641-91-2F, 5-[6-(4-Ethylphenyl)-2-oxo-1,2-dihydroindol-3ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
(2-(pyrrolidin-1-yl)ethyl)amide 342641-92-3F,
5-(6-(2,4-Dimethoxyphenyl)-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-1H-pyrrole-3-carboxylic acid
(2-(pyrrolidin-1-yl)ethyl)amide 342641-93-4F,
5-[6-(3-Isopropylphenyl)-2-oxo-1,2-dihydroindol-3ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
(2-(pyrrolidin-1-yl)ethyl)amide 342641-94-5F,
5-(5-Fluoro-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
(2-diethylaminoethyl)-3,5-dimethyl-1H-pyrrole-3-carboxylic acid
342641-95-7F, 5-(5-Dimethylulfamyl-1H-oxo-1,2-dihydroindol-3ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid
342641-96-7F, 5-(5-Dimethyl-1H-pyrrole-3-carboxylic acid
(2-(pyrrolidin-1-yl)ethyl)amide 342641-97-8F,

S-[5-(3-Chlorophenylsulfamoyl)-2-oxo-1,2-dihydroindol-3-ylidenemethyl]-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)ethyl)amide 342641-98-99, 2,4-Dimethyl-5-[2-oxo-5-(pyridin-3-ylsulfamoyl)-1,2-dihydroindol-3-ylidenemethyl]-1H-pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)ethyl)amide 342642-01-79, 5-(5-Dimethylsulfamoyl)-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 342642-02-89, 5-(5-(3-dhorophenylsulfamoyl)-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide 342642-09-59, [4-Methyl-5-(4-methyl-5-methylsulfamoyl)-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-5-methylsulfamoyl-2-oxo-1,2-dihydroindol-3-ylidenemethyl-1H-pyrrole-3-carbonyl]amino]acetic acid ethyl ester 342642-10-89,

[[4-Methyl-5-(5-methylsulfamoyl-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-1H-pyrrole-3-carbonyl]amino]acetic acid ethyl ester 342642-11-9P,

[[4-Methyl-5-(5-methylsulfamoyl-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-1H-pyrrole-3-carbonyl]amino|acetic acid 346405-32-1P, 5-(5-Fluoro-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-

Page 79 SAEED ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)ethyl) amide 35066-92-1P, 5-(5-Chloro-2-oxo-1,2-dihydroindol-3-yl)idenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-(pyrrolidin-1-yl)ethyl)amide 35066-90-1P, 5-(5-Chloro-2-oxo-1,2-dihydroindol-3-yl)idenemethyl)-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 35066-91-2P, 2,4-Dimethyl-5-(2-oxo-1,2-dihydroindol-3-yl)idenemethyl)-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl) amide 35066-92-3P, 2,4-Dimethyl-5-(2-oxo-1,2-dihydroindol-3-yl)idenemethyl)-1H-pyrrole-3-carboxylic acid (2-byrrolidin-1-ethyl) amide 35066-95-6P, 2,4-Dimethyl-5-(2-oxo-1,2-dihydroindol-3-yl)idenemethyl)-1H-pyrrole-3-carboxylic acid (2-dimethylaminoethyl) amide 35066-95-8P, 5-(5-Fluoro-2-oxo-1,2-dihydroindol-3-yl)idenemethyl)-1H-pyrrole-3-carboxylic acid (2-dimethylaminoethyl) amide 35066-95-8P, 5-(5-Fluoro-2-oxo-1,2-dihydroindol-(32)-yl)idenemethyl-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-dimethylaminoethyl) amide 35066-95-8P, 5-(5-Fluoro-2-oxo-1,2-dihydroindol-(32)-yl)idenemethyl-2,4-dimethyl-1H-pyrrole-3-carboxylic acid

dihydrolndol-(32)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic

(2-diethyl-N-oxoaminoethyl)amide 356089-03-99,
5-(5-Fluoro-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(4-methylpjerazin-1-yl)ethyl]amide 356089-04-09, 5-(5-Chloro-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(4-methylpjerazin-1-yl)ethyl]amide 356089-05-19, 5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(4-methylpjerazin-1-yl)ethyl]amide 356089-05-19, 5-(5-Bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(4-methylpjerazin-1-yl)ethyl]amide 356089-05-99, 5-(2-0xo-1,2-dihydroindol-3(32)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(3,5-dimethylpjerazin-1-yl)ethyl]amide 356089-12-0P, 5-(5-Fluoro-2-oxo-1,2-dihydroindol-(32)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(3,5-dimethylpjerazin-1-yl)ethyl]amide 356089-13-1P, 5-(5-Bromo-2-oxo-1,2-dihydroindol-(32)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(3,5-dimethylpjerazin-1-yl)ethyl]amide 356089-13-3P, 5-(5-Bromo-2-oxo-1,2-dihydroindol-(32)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(5,7-dimethyl)pjerazin-1-yl)ethyl]amide 356089-13-3P, 5-(5-Bromo-2-oxo-1,2-dihydroindol-(32)-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid [2-(5,7-dimethyl)pjerazin-1-yl)ethyl]am

(Uses)
{protein kinase modulator; prepn. of prodrugs of
 (pryrolylmethylidene)indolinones and activity as modulators of protein
 kinases)
342641-49-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-{3-(diethylamino)propyl}-2-(1-methylethyl)-N-{3-(diethylamino)propyl}-2-(1-methylethyl)-N-(9CI) (CA INDEX NAME)

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene)methyll-2-(1-methylethyll)-N-[3-(4-methyl-1-piperazinyl)propyl]-4-phenyl- (9CI) (CA INDEX NAME)

342641-54-7 CAPLUS

1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2-methyl-4-phenyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA

342641-55-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-6-(2-methoxyphenyl)-2-oxo-3H-indoi-3-ylidene|methyl|-2-methyl-4-phenyl-N-[2-(1-pyrrolidinyl)ethyl]-(9CI) (CA INDEX NAME)

L4 ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-50-3 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2-(1-methylethyl)-4-phenyl-N-{3-(1-pyrrolidinyl)propyl}-(SCI) (CA INDEX NAME)

342641-51-4 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-[2-(diethylamino)ethyl]-2-(1-methylethyl)-4-phenyl-(SCI) (CA INDEX NAME)

342641-52-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-56-9 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(dimethylamino)ethyl]-2-methyl-4-phenyl- (9CI) (CA INDEX NAME)

342641-57-0 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-(2-methoxyphenyl)-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(dimethylamino)ethyl]-2-methyl-4-phenyl-(9CI) (CA INDEX NAME)

342641-59-2 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

Page 80 SAEED ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene)methyl]-N-[3-(diethylamino)propyl]-2-methyl-4-phenyl- (9CI) (CA INDEX NAME)

342641-60-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

342641-61-6 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl}-N-{2-(dimethylamino)ethyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

342641-62-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 342641-65-0 CAPLUS LH-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[3-(1H-imidazol-1-yl))propyl]-2, 4-dimethyl- (9CI) (CA INDEX NAME)

342641-66-1 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-(2-methoxyphenyl)-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

342641-67-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-{3-methoxyphenyl}-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-63-8 CAPLUS
1H-Fyrrole-3-carboxamide, 5-[{5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-(diethylamino|ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-64-9 CAPLUS
CN lH-Fyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
342641-68-3 CAPLUS
1H-Pyrrole-3-carboxamide,
2-(diethylamino)ethyl]-5-[(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-69-4 CAPLUS
CN 1H-Fyrrole-3-carboxamide, 5-{(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-(2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX ...

342641-70-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl]-N-[3-(1H-imidazol-1-yl)propyl]-2,4-dimethyl- (9CI) (CA RNDEX NAME)

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RN 342641-71-8 CAPLUS CN 1H-Pyrrole-3-carboxamide, N-(2-(diethylamino)ethyl)-5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-72-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX

NAME)

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

RN 342641-75-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N(2-(diethylamino|ethyl)-5-[[1,2-dihydro-2-oxo-6-(3-pyridinyl)-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX

RN 342641-76-3 CAPLUS CN 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-6-(3-pyridinyl)-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX

NAME)

L4 ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-73-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl|-N-[3-(1H-imidazol-1-yl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-74-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[6-(3,5-dichlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene}methyl]-N-[2-{diethylamino}ethyl}-2,4-dimethyl- (9CI)

INDEX NAME)

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

RN 342641-77-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(1,2-dihydro-2-oxo-6-(3-pyridinyl)-3H-indol-3ylidene]methyl]-N-[3-(dimethylamino)propyl]-2,4-dimethyl- (9CI) (CA
INDEX

NAME)

342641-78-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{{1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl}-N-{3-{dimethylamino}propyl}-2,4-dimethyl-{9CI} (CA

# 17/02/2005 .

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L4 ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 342641-79-6 CAPLUS
CN 1H-Pyrcole-3-carboxamide,
N-[3-(dichylaminol)propyl]-5-[(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-80-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[3-(diethylamino)|propyl]-5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

342641-81-0 CAPLUS
1H-Indole-4-carboxamide, N-(3-chloro-4-methoxyphenyl)-3-[[4-[[3-(diethylamino)propyl]amino]carbonyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo- [9CI] (CA INDEX NAME)

NH- (CH2) 3-NEt2

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (CA INDEX NAME) (Continued)

RN 342641-85-4 CAPLUS
CN 1H-Fyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-bis(1-methylethyl)-N-[3-(1-pyrrolidinyl)propyl](9C1)

342641-87-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-(4-pyridinylmethyl)- (9CI) (CA INDEX

L4 ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-82-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-N-[3-(diethylamino)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

342641-83-2 CAPLUS lH-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl)-2,4-bis(1-methylethyl)- (9CI) (CA INDEX NAME)

342641-84-3 CAPLUS [15-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl-N-[3-(diethylamino)propyl]-2,4-bis[(1-methylethyl)- (9CI)

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-88-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[6-(4-butylphenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene[methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

342641-89-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{[1,2-dihydro-6-[2-methoxy-5-{1-

methylethyl)phenyl]-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

RN 342641-91-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(6-(4-ethylphenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene|methyl|-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- [9CI) (CA INDEX NAME)

RN 342641-92-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
[[6-(2,4-dimethoxyphenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI)
(CA INDEX NAME)

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-95-6 CAPLUS
1H-Indole-6-carboxylic acid,
4-{[[2-(dicthylamino]ethyl]amino]carbonyl
]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo- (9CI) (CA
INDEX NAME)

342641-96-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[5-[[dimethylamino]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(l-pyrrolidinyl)ethyl]-(9CI) (CA INDEX NAME)

342641-97-8 CAPLUS
1H-Pyrrole-3-carboxsmide, 5-[[5-[{(3-chlorophenyl)amino]sulfonyl]-1,2-dhydro-2-oxo-3H-indol-3-ylidenejmethyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-93-4 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{1,2-dihydro-6-[3-(1-methylethyl)phenyl]-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]-(9CI) (CA INDEX NAME)

342641-94-5 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-98-9 .CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-5-[(3-pyridinylamino)sulfonyl]-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

342642-01-7 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[[5-

[(dimethylamino)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

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342642-02-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[5-[[3-chlorophenyl)amino]sulfonyl]-1,2-dhydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

342642-09-5 CAPLUS Glycine, N-[15-[(1,2-dihydro-4-methyl-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-yildene)methyl]-4-methyl-1H-pyrrol-3-yl]carbonyl]-, ethyl ester (9CI) (CA INDEX NAME)

342642-10-8 CAPLUS Glycine, N-[5-[1,2-dihydro-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-ylidenelmethyl]-4-methyl-1H-pyrrol-3-yl]carbonyl]-, ethyl ester (9CI)

(CA INDEX NAME)

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 356068-82-1 CAPLUS
CN 1H-Pyrrole-3-catboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3yliden)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA
INDEX

356068-90-1 CAPLUS IH-Pytrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 356068-91-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino) ethyl]-5-[(1,2-dihydro-2-oxo3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342642-11-9 CAPLUS
Glycine, N-{[5-{[1,2-dihydro-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3ylidene]methyl]-4-methyl-1H-pyrrol-3-yl]carbonyl)- (9CI) (CA INDEX NAME)

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356068-92-3 CAPLUS

1H-Pyrrole-3-carboxamide,
(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

RN 356068-95-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]N-[2-(dimethylamino)ethyl)-2,4-dimethyl- [9CI) (CA INDEX NAME)

356068-96-7 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(dimethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

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356068-97-8 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-{ethylamino}ethyl]-5-[(Z)-{5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 356068-99-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-12-(idichyloxidoamino)ethyl]-5-[(z)-(5-fluoro1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA
INDEX NAME)

Double bond geometry as shown.

356069-03-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(4-methyl-1-piperazinyl)ethyl]- (9CI)(CA INDEX NAME)

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene)methyl)-2,4-dimethyl-N-[2-(4-methyl-1-piperazinyl)ethyl)- (9CI) (CA INDEX NAME)

RN 356069-07-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(1,2-dh)dydro-2-oxo-3H-indol-3-ylidene)methyl}2,4-dimethyl-N-[2-(4-methyl-1-piperezinyl)ethyl]- (9CI) (CA INDEX NAME)

356069-09-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-N-[2-(3,5-dimethyl-1-piperazinyl)ethyl)-2,4-dimethyl-

L4 ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356069-04-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(4-methyl-1-piperazinyl)ethyl)- (9CI) (CA INDEX NAME)

356069-05-1 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (9CI) (CA INDEX NAME) (Continued)

Double bond geometry as shown.

356069-12-0 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(3,5-dimethyl-1-piperazinyl)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-

(9CI)

(CA INDEX NAME)

Double bond geometry as shown.

356069-13-1 CAPLUS lH-Pyrrole-3-carboxamide, 5-{(2)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-N-[2-(3,5-dimethyl-1-piperazinyl)ethyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-15-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(3,5-dimethyl-1-piperazinyl)ethyl]-2,4-dimethyl-(SCI) (CA INDEX NAME)

Double bond geometry as shown

468745-38-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-[2-(2-pyridinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS ON STN ACCESSION NUMBER: 2002:658111 CAPLUS 137:185408

DOCUMENT NUMBER: TITLE:

3-(4-Amidopyrrol-2-ylmethylidene)-2-indolinone

INVENTOR (S):

derivatives as protein kinase inhibitors Guan, Huiping: Liang, Congxin; Sun, Li; Tang, Peng Cho; Wei, Chung Chen; Mauragis, Michael A.;

Vojkovsky,

Tomas: Jin, Qingwu: Herrinton, Paul Matthew

PATENT ASSIGNEE(S):

USA PCT Int. Appl., 167 pp. CODEN: PIXXD2

DOCUMENT TYPE: Patent English 2

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE PRIORITY APPLN. INFO.:

US 2001-312361P

P 20010815

ANSWER 55 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

3

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

(Continued)

FORMAT

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN US 2002-76140

(Continued) A3 20020215

WO 2002-US4407 W 20020215

US 2002-411732P

P 20020918

WO 2003-US4520 W 20030214

OTHER SOURCE(S):

MARPAT 137:185408

Title compds. I (R1 = H, halo, alkyl, haloalkoxy, cycloalkyl, heterocyclic, OH, alkoxy, (un)esterified CO2H, (un)substituted NH2,

R2 = H, halo, alkyl, trihalomethyl, OH, alkoxy, CN, (un)substituted NH2, SO2NH2, (un)esterified CO2H, SO2R8, R8 = alkyl, aryl, aralkyl,

heteroaryl,
heteroaryl,
heteroaryl, R3-R6 = H, alkyl; R7 = H, alkyl, aryl, heteroaryl, acyl; Z
= aryl, heteroaryl, heterocyclic, (un)substituted NH2] were prepared for

as protein kinase inhibitors in treatment of diseases, such as cancer (no data). Thus, Et 3,5-dimethyl-4-pyrrolecarboxylate was oxidized to the 5-carboxaldehyde, followed by ester hydrolysis, reaction with 5-fluoro-2-oxindole and amidation to give the amide II. 375798-55-39 452104-42-69 452104-43-7P 452104-46-89 452104-45-99 452104-46-9P 452104-46-9P 452104-46-9P 452104-48-29 452104-53-99 452104-53-99 452104-53-99 452104-53-99 452104-53-99 452104-53-99 452104-53-99 452104-53-9P 452104-53-9P

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ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
452104-56-2P 452104-57-3P 452104-68-4P
452104-55-3P 452104-60-8P 452104-61-8P
452104-65-3P 452104-60-8P 452104-67-5P
452104-66-69 452104-66-4P 452104-67-5P
452104-66-69 452104-68-7P 452104-67-5P
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452104-80-2P 452104-81-3P 452104-82-4P
452104-80-6P 452104-97-8P 452104-82-7P
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452105-19-0P 452105-20-3P 452105-21-4P
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452105-25-9P 452105-20-3P 452105-30-5P
452105-25-9P 452105-20-3P 452105-30-5P
452105-25-9P 452105-63-4P
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452105-23-9P 452105-63-4P
452105-21-9P 452105-63-4P
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452105-2105-21-9P
452105-21-9P
452105-21-9P
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452105-21-9P
452105-21-9P
4521

Double bond geometry as shown.

452104-42-6 CAPLUS 1H-Pyrrole-3-carboxamide, N-[3-{diethylamino}-2-hydroxypropyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene}methyl]-2,4-dimethyl- (9CI)(CA INDEX NAME)

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452104-45-9 CAPLUS

1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3ylidene]methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl)-2,4-dimethyl- (9CI)
(CA INDEX NAME)

452104-46-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI)
(CA INDEX NAME)

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-43-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI)(CA INDEX NAME)

RN 452104-44-8 CAPLUS
CN 1H-Pyrcole-3-carboxamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]N-(2-hydroxy-3-(4-morpholinyl)propyl)-2,4-dimethyl- (9CI) (CA INDEX

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

RN 452104-47-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl- [9CI] (CA INDEX NAME)

452104-48-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3ylidene}methyl]-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

452104-49-3 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

452104-50-6 CAPLUS

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-53-9 CAPLUS 1H-Indole-5-carboxamide, 3-{[4-{[[3-(diethylamino)-2-

hydroxypropyl]amino]carbonyl]-5-methyl-3-phenyl-1H-pyrrol-2-yl]methylene]2,3-dihydro-N-(2-hydroxyethyl)-2-oxo- (9CI) (CA INDEX NAME)

452104-54-0 CAPLUS lH-Pyrrole-3-carboxamide, N-[3-(diethylamino)-2-hydroxypropyl]-5-[[1,2-dihydro-5-(4-morpholinylcarbonyl)-2-exo-3H-indol-3-ylidene]methyl]-4-(4-fluorophenyl)-2-methyl- (9Cl) (CA INDEX NAME)

452104-55-1 CAPLUS
1H-Indole-5-carboxamide, 3-[[4-[[[3-(diethylamino)-2-hydroxypropyl]amino]carbonyl]-3-(4-fluorophenyl)-5-methyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-N-(1-methylethyl)-2-oxo- (9CI) (CA INDEX NAME)

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

452104-51-7 CAPLUS 1H-Indole-5-carboxamide, 3-{[4-[{[3-(diethylamino)-2-

hydroxypropyl]amino]carbonyl]-5-methyl-3-phenyl-1H-pyrrol-2-yl]methylene]2,3-dihydro-2-oxo-N-phenyl- (9CI) (CA INDEX NAME)

452104-52-8 CAPLUS 1H-Indole-5-carboxamide, 3-[[4-[[[3-(diethylamino)-2-

hydroxypropyl]amino]carbonyl]-5-methyl-3-phenyl-1H-pyrrol-2-yl]methylene}2,3-dihydro-N-methyl-2-oxo- (9CI) (CA INDEX NAME)

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-56-2 CAPLUS 1H-Indole-5-carboxamide, 3-[[4-[[[3-(diethylamino)-2-

hydroxypropyl|amino]carbonyl]-3-(2,4-difluorophenyl)-5-methyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-N-phenyl- (9CI) (CA INDEX NAME)

452104-57-3 CAPLUS 1H-Indole-5-carboxamide, 3-[{4-[[[3-(diethylamino)-2-

hydroxypropyl]amino]carbonyl]-3-(2,4-difluorophenyl)-5-methyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-N-(2-hydroxyethyl)-2-oxo-(9CI) (CA INDEX NAME)

452104-58-4 CAPLUS

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
1H-Indole-5-carboxamide, 3-[[3-(4-cyanophenyl)-4-[([3-(diethylamino)-2-hydroxypropyl]amino]carbonyl]-5-methyl-1H-pyrrol-2-yl]methylene]-2,3dihydro-N,N-dimethyl-2-oxo- (9CI) (CA INDEX NAME)

452104-59-5 CAPLUS 1H-Pyrrole-3-carboxamide, 4-(4-cyanophenyl)-N-[3-(diethylamino)-2-

hydroxypropyl)-5-[[1,2-dihydro-5-(4-morpholinylcarbonyl)-2-oxo-3H-indol-3-ylidene]methyl]-2-methyl- (9CI) (CA INDEX NAME)

452104-60-8 CAPLUS 1H-Indole-5-carboxamide, 3-{[3-(4-chlorophenyl)-4-[{[3-(diethylamino)-2-hydroxypropyl]amino]carbonyl]-5-methyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-N-phenyl- (9CI) (CA INDEX NAME)

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-61-9 CAPLUS
1H-Indole-5-carboxamide, 3-[(3-(4-chlorophenyl)-4-[(3-(diethylamino)-2-hydroxypropyl]amino)carbonyl]-5-methyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-N-(1-methylethyl)-2-oxo- (9CI) (CA INDEX NAME)

452104-62-0 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-hydroxy-3-(2H-tetrazol-2-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-63-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-(2-hydroxy-3-(2H-tetrazol-2-yl)propyl]-2,4-dimethyl-(SCI) (CA INDEX NAME)

452104-64-2 CAPLUS IH-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-5-(trifluoromethoxy)-3H-indoi-3-ylidene]methyl]-N-{2-hydroxy-3-(2H-tetrazol-2-yl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

(Continued)

452104-65-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(1H-tetrazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

452104-66-4 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-(2-hydroxy-3-(1H-tetrazol-1-yl)propyl)-2,4-dimethyl-9CI) (CA INDEX NAME)

452104-67-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-5-(trifluoromethoxy)-3H-indol-3-ylidene]methyl]-N-[2-hydroxy-3-(lH-tetrazol-1-yl)propyl]-2,4-dimethyl- (9C1) (CA INDEX NAME)

452104-68-6 CAPLUS
1H-Pyrrole-3-carboxamide, N-[3-(2,6-dimethyl-4-morpholinyl)-2-

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-70-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-5-(trifluoromethoxy)-3H-

indol-3-ylidene]methyl]-N-[3-(2,6-dimethyl-4-morpholinyl)-2-hydroxypropyl]2,4-dimethyl- (9CI) (CA INDEX NAME)

452104-71-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-((5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl|-N-(2-hydroxy-3-(3-methyl-2,5-dioxo-1-imidazolidinyl)propyl|-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
hydroxypropyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]2,4-dimethyl- (9CI) (CA INDEX NAME)

452104-69-7 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[{5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[3-(2,6-dimethyl-4-morpholinyl)-2-hydroxypropyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-72-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-5-(trifluoromethoxy)-3H-indol-3-ylidene]methyl]-N-[2-hydroxy-3-(3-methyl-2,5-dioxo-1-imidazolidinyl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

452104-73-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(3-methyl-2,5-dioxo-1-imidazolidinyl)propyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

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RN 452104-74-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(1,2-dihydro-2-xox-3H-indol-3-ylidene)methyl]N-[3-(1,1-dioxido-4-thiomorpholinyl)-2-hydroxypropyl]-2,4-dimethyl- (9CI)
(CA INDEX NAME)

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-76-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene) methyl] -N- (3-(1,1-dioxido-4-thiomorpholinyl) -2-hydroxypropyl]-2,4dimethyl- (9CI) (CA INDEX NAME)

452104-77-7 CAPLUS

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-75-5 CAPLUS
1H-Pyrrole-3-carboxamide, N-[3-{1,1-dioxido-4-thiomorpholiny1}-2-hydroxypropy1]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methy1}-2,4-dimethy1- (9CI) (CA INDEX NAME)

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 1H-Pyrrole-3-carboxamide, 5-((5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[3-{1,1-dioxido-4-thiomorpholinyl)-2-hydroxypropyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

452104-78-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-N,2,4-trimethyl-(9CI) (CA INDEX NAME)

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- L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
- 452104-79-9 CAPLUS
  1H-Pyrrole-3-carboxamide, 5-([5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-hydroxy-3-(3H-1,2,3-triazolo[4,5-b]pyridin-3-yloxy)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

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- 452104-80-2 CAPLUS

  IH-Pyrrole-3-carboxamide, 5-{(5-chloro-1,2-dihydro-2-oxo-3H-indol-3ylidene]methyl]-M-(2-hydroxy-3-(3H-1,2,3-triazolo(4,5-b)pyridin-3yloxy)propyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)
- L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

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(Continued)

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- 452104-82-4 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-
- ylidene)methyl}-N-[2-hydroxy-3-[(3-oxido-1H-benzotriazol-1-yl)oxy]propyl}2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

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- RN 452104-81-3 CAPLUS CN 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-5-(trifluoromethoxy)-3H-
- indol-3-ylidene|methyl]-N-[2-hydroxy-3-(3H-1,2,3-triazolo[4,5-b]pyridin-3-yloxy)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

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- RN 452104-83-5 CAPLUS
  CN 1H-Pyrrole-3-carboxamide, 5-{(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-
- ylidene)methyl}-N-[2-hydroxy-3-[(3-oxido-1H-benzotriezol+1-yl)oxy)propyl]2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

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452104-84-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{[1,2-dihydro-2-oxo-5-(trifluoromethoxy)-3H-indol-3-ylidene]methyl]-N-{2-hydroxy-3-{(3-oxido-1H-benzotriazol-1-yl)oxy]propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

(Continued)

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 $452104-85-7 \quad CAPLUS \\ 1H-Pyrrole-3-carboxamide, 5-[{z}]-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2, 4-dimethyl- (9CI) (CA INDEX NAME)$ 

Double bond geometry as shown.

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-86-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

452104-87-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-{2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl- (9CI)(CA INDEX NAME)

Double bond geometry as shown.

452104-88-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI)(CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-89-1 CAPLUS 1H-Pyrrole-3-carboxamide, 5-((Z)-(1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl)-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-90-4 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as sho

452104-91-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl}-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-92-6 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-93-7 CAPLUS 1H-Indole-5-carboxamide, 3-[[4-[[[3-(diethylamino)-2-

hydroxypropyl]amino]carbonyl]-5-methyl-3-phenyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-N-phenyl-, (32)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-94-8 CAPLUS 1H-Indole-5-carboxamide, 3-[[4-[[[3-(diethylamino)-2-

hydroxypropyl]amino]carbonyl]-5-methyl-3-phenyl-1H-pyrrol-2-yl]methylene)-2,3-dihydro-N-methyl-2-oxo-, (3Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) hydroxypropyllamino|carbonyl]-3-(4-fluorophenyl)-5-methyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-N-(1-methylethyl)-2-oxo-, (32)- (9CI) (CA INDEX

Double bond geometry as shown.

452104-98-2 CAPLUS 1H-Indole-5-carboxamide, 3-[[4-[[[3-(diethylamino)-2-

hydroxypropyl]amino]carbonyl]-3-{2,4-difluorophenyl}-5-methyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-N-phenyl-, (3Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-99-3 CAPLUS 1H-Indole-5-carboxamide, 3-[[4-[[[3-(diethylamino)-2-

hydroxypropyl]amino|carbonyl]-3-(2,4-difluorophenyl)-5-methyl-1H-pyrrol-2-yl|methylene)-2,3-dihydro-N-(2-hydroxyethyl)-2-oxo-, (3Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452104-95-9 CAPLUS 1H-Indole-5-carboxamide, 3-{[4-[[{3-(diethylamino)-2-

hydroxypropyl amino carbonyl - 5-methyl - 3-phenyl - 1H-pyrrol - 2-yl methylene - 2,3-dihydro-N-(2-hydroxyethyl) - 2-oxo-, (3Z) - (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 452104-96-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[3-(diethylamino)-2-hydroxypropyl]-5-[(Z)-[1,2-dihydro-5-(4-morpholinylcarbonyl)-2-oxo-3H-indol-3-ylidene]methyl]-4-(4-fluorophenyl)-2-methyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

452104-97-1 CAPLUS
1H-Indole-5-carboxamide, 3-[[4-[[[3-(diethylamino)-2-

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452105-00-9 CAPLUS 1H-Indole-5-carboxamide, 3-[[3-(4-cyanophenyl)-4-[[3-(diethylamino)-2-hydroxypropyl]amino]carbonyl]-5-methyl-1H-pyrrol-2-yl]methylene]-2,3-dlhydro-N,N-dlmethyl-2-oxo-, (32)- (921) (CA INDEX NAME)

Double bond geometry as shown.

452105-01-0 CAPLUS 1H-Pyrrole-3-carboxamide, 4-(4-cyanophenyl)-N-[3-(diethylamino)-2-hydroxypropyl]-5-[(2)-[1,2-dihydro-5-(4-morpholinylcarbonyl)-2-oxo-3H-indol-3-ylidene)methyl]-2-methyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 452105-02-1 CAPLUS 1H-Indole-5-carboxamide, 3-[[3-(4-chlorophenyl)-4-[[[3-(diethylamino)-2-hydroxypropyl]amino]carbonyl]-5-methyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-N-phenyl-, (3Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

452105-03-2 CAPLUS
1H-Indole-5-carboxamide, 3-[[3-(4-chlorophenyl)-4-[[[3-(diethylamino)-2-hydroxypropyl]amino]carbonyl]-5-methyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-N-(1-methylethyl)-2-oxo-, (32)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

452105-04-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-N-[2-hydroxy-3-(2H-tetrazol-2-yl)propyl)-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452105-08-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(1H-tetrazol-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as sho

452105-09-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{{Z}-{1,2-dihydro-2-oxo-5-{trifluoromethoxy}-3-dih-y-2-dihydro-2-oxo-5-{trifluoromethoxy}-3-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

452105-10-1 CAPLUS
1H-Pyrrole-3-carboxamide, N-[3-[(2R,6S)-2,6-dimethyl-4-morpholinyl]-2-hydroxypropyl]-5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl-, rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.
Double bond geometry as shown.

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\$$

452105-05-4 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(Z)-{5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(2H-tetrazol-2-yl)propyl}-2,4-dimethyl-(SCI) (CA INDEX NAME)

Double bond geometry as shown.

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & &$$

Double bond geometry as shown.

452105-07-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{Z}]-{5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-(1H-tetrazol-1-yl)propyl}-2,4-dimethyl-(SCI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452105-11-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[3-[(2R,6S)-2,6-dimethyl-4-morpholinyl]-2-hydroxypropyl}2,4-dimethyl-, rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.
Double bond geometry as shown

 $\begin{array}{lll} 452105-12-3 & CAPLUS \\ 1H-Pyrrole-3-carboxamide, & 5-[(Z)-[1,2-dihydro-2-oxo-5-(trifluoromethoxy)-3H-indo-3-ylidene]methyl]-N-[3-[(2R,65)-2,6-dimethyl-4-morpholinyl]-2-hydroxypropyl]-2,4-dimethyl-, rel- (9CI) & (CA INDEX NAME) \\ \end{array}$ 

Relative stereochemistry.
Double bond geometry as shown.

452105-13-4 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-H-[(2R)-2-hydroxy-3-(3-methyl-2,5-dloxo-1-imidazolidinyl)propyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

452105-14-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-H-[(2R)-2-hydroxy-3-(3-methyl-2,5-dioxo-1-imidazolidinyl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

452105-15-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-[1,2-dihydro-2-oxo-5-(trifluoromethoxy)-3H-indo-3-ylidene|methyl]-M-[(ZR)-2-hydroxy-3-(3-methyl-2,5-dioxo-1-imidazolidinyl)propyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

452105-16-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-{(2S)-2-hydroxy-3-(3-methyl-2,5-dloxo-1-imidazolidinyl)propyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

(Continued) ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

452105-20-3 CAPLUS
1H-Pyrrole-3-carboxamide, N-[3-(1,1-dioxido-4-thiomorpholinyl)-2-hydroxypropyl]-5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

452105-21-4 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[3-(1,1-dioxido-4-thiomorpholinyl)-2-hydroxypropyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

452105-22-5 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(2)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[3-(1,1-dioxido-4-thiomorpholinyl)-2-hydroxypropyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

Absolute stereochemistry. Double bond geometry as shown.

452105-18-9 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(2)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[(28)-2-hydroxy-3-(3-methyl-2,5-dloxo-1-imidazolidinyl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

452105-19-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-{3-{1,1-dioxido-4-thiomorpholinyl}-2-hydroxypropyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452105-23-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-((28)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown

452105-24-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-M-{(2R)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

452105-25-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-((ZB)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

452105-26-9 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-{(ZS)-2-hydroxy-3-(4-morpholinyl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

452105-27-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-(2-hydroxy-3-(3H-1,2,3-triazolo[4,5-b]pyridin-3-yloxy)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

452105-28-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[(2-hydroxy-3-(3H-1,2,3-triazolo[4,5-b]pyridin-3-yloxy)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452105-31-6 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl)-N-{2-hydroxy-3-{(3-oxido-1H-benzotriazol-1-yl)oxy}propyl}2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

452105-32-7 CAPLUS  $\begin{array}{lll} & & & & \\$ 

Double bond geometry as shown.

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

452105-29-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{Z}-[1,2-dihydro-2-oxo-5-(trifluoromethoxy)-

3H-indol-3-ylidene]methyl]-N-[2-hydroxy-3-(3H-1,2,3-triazolo[4,5-b]pyridin-3-yloxy)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

PAGE 1-B

452105-30-5 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[2-hydroxy-3-[(3-oxido-1H-benzotriazol-1-yl)oxy]propyl]2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

PAGE 1-B



452105-44-1 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(2)-(1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methy1]-N-[{2S}-3-(1,1-dioxido-4-thiomorpholiny1)-2-hydroxypropy1]2,4-dimethy1- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

Absolute stereochemistry.
Double bond geometry as shown.

452105-46-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methy1)-N-{(2S}-3-(1,1-dioxido-4-thiomorpholiny1)-2-hydroxypropy1}2,4-dimethy1-(9CI) (CA INDEX NAME)

Absolute stereochemistry.

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L4 ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN Double bond geometry as shown. (Continued)

452105-47-4 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{{Z}}-{5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[(25)-3-(1,1-dioxido-4-thiomorpholinyl)-2-hydroxypropyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

452105-62-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[(2S)-2-hydroxy-3-(4-morpholinyl)propyl]-N,2,4-trimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

452105-63-4 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl]-N-[(2R)-2-hydroxy-3-(4-morpholinyl)propyl]-N,2,4-trimethyl-

L4 ANSWER 57 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN ACCESSION NUMBER: 2002:541364 CAPLUS DOCUMENT NUMBER: 138:147266 Inhibition of Cartesian Control of Cartesian Capture Capt

138:147266
Inhibition of constitutively active forms of mutant kit by multitargeted indolinone tyrosine kinase inhibitors
Liao, Albert T.; Chien, May B.; Shenoy, Narmada; Mendel, Dirk B.; McMahon, Gerald; Cherrington, Julie M.; London, Cheryl A.
Department of Surgical and Radiological Sciences, School of Veterinary Medicine, University of California at Davis, Davis, CA, 95616, USA Blood (2002), 100(2), 585-593
CODEN: BLOOAW; ISSN: 0006-4971
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Mutations in the proto-oncogene c-kit, including point mutations,

deletions, or duplications in the neg. regulatory juxtamembrane (JM)

domain or point mutations in the catalytic domain, have been observed in

human and canine cancers and often result in constitutive activation of

Kit in the absence of ligand binding. To identify a receptor tyrosine

kinase (RTK) inhibitor capable of blocking the function of mutant Kit, we

evaluated 3 indolinones (SU11652, SU11654, and SU11655) that act as

competitive inhibitors of ATP binding to several members of the split

kinase family of RTKs, including VEOFR, FGFR, PGFR, AGK Att. Mast cell

lines expressing either wildtype (WT) Kit, a point mutation in the JM

domain, a tandem duplication in the JM domain, or a point mutation in the

catalytic domain were used for these studies. All 3 indolinones

inhibited

bited phosphorylation of WT Kit in the presence of stem cell factor at concns. as low as 0.01 µM. Autophosphorylation of both JM mutants was inhibited at 0.01 to 0.1 µM, resulting in cell cycle arrest within 24 h, whereas autophosphorylation of the catalytic domain mutant was inhibited at 0.25 to 0.5 µM, resulting in cell death within 24 h. Poly(ADP-ribose) polymerase (PARP) cleavage was noted in all Kit mutant lines after indolinone treatment. In summary, SU1652, SU11654, and SU11655 are effective RTK inhibitors capable of disrupting the function

all forms of mutant Kit. Because the concns. of drug necessary for receptor inhibition are readily achievable and nontoxic in vivo, these compds. may be useful in the treatment of spontaneous cancers expressing kit mutations.

326914-10-7, SU 11652 356068-94-5, SU 11654

356069-35-7, SU 11655

RL: PRC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(inhibition of constitutively active forms of mutant kit by multitargeted indollance tyrosine kinase inhibitors)

326914-10-7 CAPLUS

1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene) methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 56 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (9CI) (CA INDEX NAME) (Continued)

Absolute stereochemistry. Double bond geometry as shown

REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

ANSWER 57 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

356068-94-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX

Double bond geometry as shown.

356069-35-7 CAPLUS H-Pyrrole-3-carboxamide, 5-[(2)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-{2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

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17/02/2005

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L4 ANSWER 57 OF 65 CAPLUS COPYRIGHT 2005 ACS ON STN (Continued) REFERENCE COUNT: 87 THERE ARE 87 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE

137:109202
Preparation of 4-aryl substituted indolinones as protein kinase signal transduction modulators for inhibiting abnormal cell proliferation Cui, Jingrahnormal cell proliferation Cui, Jingrahnormal cell Shen, Hong; Chu, Ji Yu; Zhang, Fang-Jie; Koenig, Marcel; Do, Steven Huy; Li, Xiaoyuan; Wei, Chung Chen; Tang, Peng Cho USA
PCT Int. Appl., 560 pp.
CODEN: PIXXD2
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English
1 INVENTOR (S): PATENT ASSIGNEE(S): SOURCE: DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO.

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W: AE, AG, AL, AM, AT, AU, AR, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, NN, MW, MW, MZ, NO, MZ, OM, PL, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZM, ZM, ZM, AZ, BY, KG, KZ, MD, TT, TJ, TM

RW: GM, GM, KE, LS, MM, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GG, GW, ML, MR, NS, NT, DT, GC A2432114

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A1 20030410

EP 2001-23488

A2 20011220

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FR, OM, CY, AL, TR

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T2 20040013

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FR, OM, KC, YA, AL, TR

JP 2004518669

T2 20040013

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FR, OM, KC, YA, AL, TR

JP 2004518669

T2 20040012

US 2001-23488

A3 20011220

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L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN ACCESSION NUMBER: 2002:539677 CAPLUS DOCUMENT NUMBER: 137:109202

TITLE:

ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

Title compds. I [Rl = {un}substituted aryl or heteroaryl: R2 = H, halo, alkyl, alkenyl, alkynyl, heterocyclyl, etc.: R3 = {un}substituted pyrrole or cycloalkenylpyrrole, as well as pharmaceutical compns. thereof, are

RN 442558-07-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino|ethyl]-5-[[4-(4-fluorophenyl)1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA
INDEX NAME)

442558-08-9 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[4-(4-fluorophenyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

442558-09-0 CAPLUS

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SAEED

ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) prepd. and disclosed as compds. capable of modulating protein kinase signal transduction in order to regulate, modulate and/or inhibit ANSWER 58 07 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) prepd. and disclosed as compale of modulate and/or inhibit ormal cell proliferation. Thus II, was prepd. via content and/or inhibit ormal cell proliferation. Thus II, was prepd. via contentation of 4-phenyl-1, 3-dihydroindol-2-one with 5-formyl-2-methyl-4-{3-(4-methylpiperazin-1-yl)propyl]-1H-pyrrole-3-carboxylic acid Et ester. I were evaluated against eight specfic kinases, e.g., FGFRI, for which I possessed ICSO values (µM) of 0.0091-2.07. The present invention also relates to methods for treating protein kinases related disorders. 442558-05-69 442558-15-89 442558-26-99 442558-20-99 442558-15-99 442558-19-99 442558-21-99 442558-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 442568-31-99 44 (target compound; preparation of (aryl) (pyrrolylmethylene) indolinones protein kinase signal transduction modulators)
442558-05-6 CAPLUS
1H-Pyrrole-3-carboxamide,
-(dlethylamino)ethyl]-5-[(1,2-dihydro-2-oxo4-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME) ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
1H-Pyrrole-3-carboxamide, 5-[[4-(4-fluorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene|methyl-2,4-dimethyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl](9CI) (CA INDEX NAME)

RN 442558-15-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-[[4-(3-fluorophenyl)1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- [9CI) (CA INDEX NAME)

442558-16-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI)
(CA INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

442558-24-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[4-(4-chlorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

442558-25-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[4-(4-chlorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl]-(9CI) (CA INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

442558-17-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl]-(9CI) (CA INDEX NAME)

442558-23-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[4-(4-chlorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI)

INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

442558-31-8 CAPLUS IH-Pyrrole-3-carboxamide, 5-[[4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI)

442558-32-9 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

RN 442558-33-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{{4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl}-2,4-dimethyl-N-{2-(1H-1,2,3-triazol-1-yl)ethyl}-(9CI) (CA INDEX NAME)

RN 442558-39-6 CAPLUS

(N 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[[1,2-dihydro-4-(4-methoxyphenyl)-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

RN 442558-40-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{[1,2-dihydro-4-(4-methoxypheny1)-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidiny1)ethyl]- (9CI) (CA INDEX NAME)

(Continued)

RN 442558-41-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-4-(4-methoxyphenyl)-2-oxo-3H-indol-3-ylidene|methyl}-2,4-dimethyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl)-(9CI) (CA INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442558-48-7 CAPLUS
IN-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl)-5-[[1,2-dihydro-4-(3-methoxyphenyl)-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 442558-49-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-4-(3-methoxyphenyl)-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442558-50-1 CAPLUS

CN 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-4-(3-methoxyphenyl)-2-oxo-3H-indol-3-ylidenelmethyl]-2,4-dimethyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl](9C1) (CA INDEX NAME)

RN 442558-54-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide, N-[2-[bis(1-methylethyl)amino]ethyl]-5-[[1,2-dihydro-4-(3-methoxyphenyl)-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

RN 442558-56-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[4-(4-bromophenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene)methyl]-N-[2-(diethylamino)ethyl)-2,4-dimethyl- (9CI) (CA INDEX
NAME\*

RN 442558-57-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[4-(4-bromophenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene|methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 44258-63-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
[[4-(4-bromophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-4-methyl- (9CI) (CA INDEX NAME)

RN 442558-65-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(4-(3-bromophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl-(9CI) (CA INDEX
NAME)

RN 442558-66-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[4-(3-bromophenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene|methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl}- (9CI) (CA INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442558-58-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[{4-(4-bromophenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene|methyl}-2,4-dimethyl-N-{2-(1H-1,2,3-triazol-1-yl)ethyl}- (9CI)
(CA INDEX NAME)

RN 442558-62-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide, N-{2-[bis{1-methylethyl}amino]ethyl]-5-[[4-(4-bromophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-(9C1) (CA INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442558-67-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[4-(3-bromophenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene]methyl]-2,4-dimethyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl]- (9CI)
(CA INDEX NAME)

RN 442558-71-6 CAPLUS

N 1H-Pyrrole-3-carboxamide, N-[2-[bis(1-methylethyl)amino]ethyl]-5-[[4-(3-bromophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

NH-CH2-CH2-N(Pr-1)2

RN 442558-88-5 CAPLUS

(N 1H-Pyrrole-3-carboxamide,
N-[2-(idethylamino)=thyl]-5-[[4-(4-fluorophenyl)1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-4-methyl- (9CI) (CA INDEX NAME)

CH2-CH2-NEt2

RN 442558-89-6 CAPLUS
CN 1H-Pytrole-3-carboxamide,
N-[2-(diethylamino)=thyl]-5-[[4-(3-fluorophenyl)1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-4-methyl- (9CI) (CA INDEX NAKE)

NH-CH2-CH2-NEt2

442558-90-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[4-(4-chloropheny1)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-4-methyl- (9CI) (CA INDEX NAME)

ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN NAME)

CH2-CH2-NEt2

442558-94-3 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl)-5-[[1,2-dihydro-4-(3-methoxyphenyl)-2-oxo-3H-indol-3-ylidene]methyl)-4-methyl- (9CI) (CA

- CH2- CH2- NEt2

RN 442558-96-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(idethylamino|ethyl)-5-[[4-(2-fluorophenyl)1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA
INDEX NAME)

NH-CH2-CH2-NEt2

442558-97-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

CH2-CH2-NEt2

442558-91-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-4-methyl- (9CI) (CA INDEX NAME)

RN 442558-92-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(4-3-bromophenyl)-1-,2-dihydro-2-oxo-3H-indol3-ylidene|methyl]-N-[2-(diethylamino)ethyl]-4-methyl- (9CI) (CA INDEX NAME)

NH-CH2-CH2-NEt2

442558-93-2 CAPLUS 1H-Pyrcole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(1,2-dihydro-4-(4-methoxyphenyl)-2-oxo-3H-indol-3-ylidene|methyl]-4-methyl- (9CI) (CA

ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

442558-98-7 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[4-[2-fluoropheny1]-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2, 4-dimethyl-N-[2-[1H-1,2,3-triazol-1-yl)ethyl]-(9CI) (CA INDEX NAME)

442559-03-7 CAPLUS 1H-Pyrrole-3-carboxamide, N-[3-{diethylamino}propyl}-5-[{4-{4-fluorophenyl}-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

NH- (CH2) 3-NEt2

442559-05-9 CAPLUS
1H-Pyrrole-3-carboxamide, N-[3-(diethylamino)propyl]-5-[{4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl)-2,4-dimethyl-(3C1) (CA INDEX RAME)

n- (CH<sub>2</sub>) 3-NEt<sub>2</sub>

RN 442559-07-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-[[1,2-dihydro-2-oxo-4-[3-(trifluoromethyl)phenyl]-3H-indol-3-ylidene]methyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

CH2-CH2-NEt2

442559-08-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-4-[3-(trifluoromethyl]phenyl]-3H-indol-3-ylidene]methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

NH- (CH2) 3-NEt2

442559-19-5 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[4-(3-chloro-4-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

ľ NH-CH2-CH2-NEt2

442559-20-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[4-(3-chloro-4-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442559-09-3 CAPLUS CN 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-4-[3-

(trifluoromethyl)phenyl]-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl]- (9CI) (CA INDEX NAME)

RN 442559-14-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[3-(diethylamino)propyl]-5-[[1,2-dihydro-2-oxo4-[3-(trifluoromethyl)phenyl]-3H-indol-3-ylidene]methyl]-2,4-dimethyl{9CI} (CA INDEX NAME)

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442559-42-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)ethyl]-5-{[1,2-dihydro-2-oxo-4-(4-(trifluoromethoxy)phenyl]-3H-indol-3-ylidene)methyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

442559-43-5 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-4-[4-

(trifluoromethoxy)phenyl]-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl]- (9CI) (CA INDEX NAME)

RN 442559-53-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide, N-[3-(diethylamino)propyl]-5-[(2)-[4-(4fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-4-methyl(9CI)
(CA INDEX NAME)

Double bond geometry as shown.

RN 442559-54-8 CAPLUS
CN 1H-Pytrole-3-carboxamide,
5-[(2)-[(4-3-chloropheny1)-1,2-dihydro-2-oxo-3Hindol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-4-methyl- (9CI) (CA
INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442559-79-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[{Z}-[1,2-dihydro-4-[3-{2-hydroxyethyl)phenyl}-

2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl](9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 442559-80-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[1,2-dihydro-4-]3-(2-hydroxyethyl)phenyl}2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-{2-(1H-1,2,3-triazol-1-yl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442559-55-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene]methyl]-N,2,4-trimethyl-N-(1-methyl-4-piperidinyl)- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

RN 442559-57-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-{{Z}}-{{4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442559-97-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(3)-(4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene)methyl]-4-methyl-N-[2-(4-morpholinyl)ethyl]- (9CI) (CA
INDEX NAME)

Double bond geometry as shown.

RN 442559-98-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-{(2)-{4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene|methyl)-4-methyl-N-{3-(2-methyl-1-piperidinyl)propyl}(9CI) {CA INDEX NAME}

Double bond geometry as shown.

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RN 442559-99-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-(4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene]methyl]-4-methyl-N-[2-(1-piperidinyl)ethyl]- (9CI) (CA
INDEX NAME)

Double bond geometry as shown.

RN 442560-00-1 CAPLUS
CN 1H-Pytrole-3-carboxamide,
5-[(2)-[(4-[3-chlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene]methyl]-4-methyl-N-[2-(4-pyridinyl)ethyl]- (9CI) (CA
INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) indol-3-ylidene|methyl|-4-methyl-N-[3-(2-oxo-1-pyrrolidinyl)propyl]-(CA INDEX NAME)

Double bond geometry as shown.

RN 442560-03-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-{(2)-(4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene|methyl|-N-{2-hydroxy-3-(4-morpholinyl)propyl}-4-methyl(9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 442560-04-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-{(2)-(4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene]methyl]-N-(2-hydroxy-3-(1H-1,2,3-triazol-1-yl)propyl]-4methyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442560-01-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-{(Z)-(4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3H-indo-3-ylidene]methyl]-4-methyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 442560-02-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(Z)-[4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3H-

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442560-07-8 CAPLUS

IH-Pyrrole-3-acetamide, 5-{(Z)-{4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl)-2,4-dimethyl-N-{2-(4-morpholinyl)ethyl}- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 442560-09-0 CAPLUS
CN 1H-Pyrrole-3-acetamide, 5-[(Z)-[4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-yliden@methyl]-2,4-dimethyl-N-[2-(1-piperidinyl)ethyl]- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

442560-10-3 CAPLUS H-Pyrrole-3-acetamide, 5-[(2)-[4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(4-pyridinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

442560-11-4 CAPLUS  $\begin{array}{lll} & \text{CAPLUS} \\ & \text{H-Pyrrole-3-acetamide,} & & \text{5-[\{Z\}-[4-\{2-fluorophenyl\}-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) \\ & \text{(CA INDEX NAME)} \\ \end{array}$ 

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

442560-25-0 CAPLUS
1H-Pyrrole-3-acetamide, 5-[(Z)-[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-yiidene]methyl)-2,4-dimethyl-N-[2-(4-morpholinyl)ethyl]- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

442560-26-1 CAPLUS
1H-Pyrrole-3-acetamide, N-[2-(diethylamino)ethyl]-5-[(2)-[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

442560-14-7 CAPLUS 1H-Pyrrole-3-acetamide, N-[2-(diethylamino)ethyl]-5-[(Z)-[4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

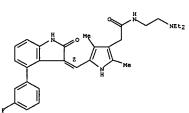
Double bond geometry as shown.

442560-22-7 CAPLUS 1H-Pyrrole-3-acetamide, 5-[(Z)-[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-

indol-3-ylidene)methyl]-N-{2-hydroxy-3-(2H-1,2,3-triazol-2-yl)propyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN



442560-27-2 CAPLUS
1H-Pyrrole-3-acetamide, 5-[(Z)-[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidenejmethyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

442560-29-4 CAPLUS 1H-Pyrrole-3-acetamide, 5-[(2)-[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(4-pyridinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 442560-39-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-{(2)-(4-{1,1'-biphenyl}-3-yl-1,2-dihydro-2-oxo3H-indol-3-ylidene)methyl]-N-{2-hydroxy-3-(2H-1,2,3-triazol-2-yl)propyl}2,4-dimethyl- (9CI) {CA INDEX NAME}

Double bond geometry as shown.

RN 442560-43-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-(25)-(4-3-fluorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidenejmethyl]-2,4-dimethyl-N-[2-(1-piperidinyl)ethyl]- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442560-64-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene|methyl]-N-methyl-N-(1-methyl-4-piperidinyl)-4-phenyl(9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 442560-65-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-[4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-4-phenyl(9CI)
(CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442560-44-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2]-(4-{1,1'-biphenyl]-2-yl-1,2-dihydro-2-oxo3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

RN 442560-51-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-{(2)-{4-(3,5-difluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl-N-[2-{1-pyrrolidinyl}ethyl}(9C1)
(CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442560-66-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-4-phenyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 442560-69-2 CAPLUS
CN lH-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(2)-[4-[3-{2-(dimethylamino)-2-oxoethyl]phenyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- [9CI) (CA INDEX NAME)

RN 442560-79-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(Z)-[4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene|methyl]-2,4-dimethyl-N-[(1,2,3,6-tetrahydro-2,6-dioxo-4pyrimidinyl)methyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 442560-80-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2]-[4-(3-amino-1H-indazol-5-yl)-1,2-dihydro2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl(9CI) (CA INDEX NAME)

Double bond geometry as shown.

14 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442560-88-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-([2]-[4-[4-dinucropheny1]-1,2-dihydro-2-oxo3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl](9CI)
(CA INDEX NAME)

Double bond geometry as shown.

RN 442561-01-5 CAPLUS
CN 1H-Pytrole-3-carboxamide,
5-[(2),[4-(2,6-dif[luorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl](9CI)
(CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442560-81-8 CAPLUS CN 1H-Pyrrole-3-carboxamide, 5-[(Z)-[4-(3-amino-1H-indazol-5-yl)-1,2-dihydro-

2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl](9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 442560-83-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-hydroxy-3-(1-pyrrolidinyl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442561-06-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-[4-acety]-1-piperazinyl)ethyl]-5-[[Z]-[4-[2fluorophenyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-4-methyl(SCI)

Double bond geometry as shown.

RN 442561-07-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2]-[4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene]methyl]-4-methyl-N-[2-(1-piperidinyl)ethyl]- [9CI] (CA INDEX NAME)

Double bond geometry as shown.

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SAEED

RN 442561-11-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[4-3-chlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene]methyl]-N-[2-hydroxy-3-(1-pyrrolidinyl)propyl]-4-methyl(SCI) (CA INDEX NAME)

Double bond geometry as shown.

RN 442561-12-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)=(4-(2-fluoropheny1)-1,2-dihydro-2-oxo-3Hindol-3-ylidene|methyl]-N-[(2-hydroxy-3-(1-pyrrolidiny1)propy1)-4-methyl(SCI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

442561-48-0 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(Z)-{4-(3,4-dimethoxyphenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl-N-{2-(1-pyrrolidinyl)ethyl}-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

442561-52-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-[1,2-dihydro-4-[3-[(methylamino)carbonyl]phenyl]-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442561-13-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-(12)-(4-3-chlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene|methyl]-N-(2-hydroxy-3-(1-pyrrolidinyl)propyl)-2,4dimethyl- (9C1) (CA IMDEX NAME)

Double bond geometry as shown.

 $\begin{array}{lll} 442561-39-9 & \text{CAPLUS} \\ 1\text{H-Pyrrole-3-carboxamide,} & 5-\left[(z)-\left[1,2-\text{dihydro-4-}\left[3-\left[2-\text{hydroxyethoxy}\right]\right]-2-\text{oxo-3H-indol-3-ylidene}\right] \text{methyl}-2, 4-\text{dimethyl-N-}\left[2-\text{hydrolidinyl}\right] \text{ethyl}- & (\text{GA INDEX NAME}) \end{array}$ 

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

RN 442561-53-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-{(2)-(4-(4-chlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene)methyl]-N-[2-(1,1-dioxido-4-thiomorpholinyl)ethyl}-2,4dimethyl- {9CI} (CA INDEX NAME)

Double bond geometry as shown.

442561-54-8 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(1,1-dioxido-4-thiomorpholinyl)ethyl]-5[(2)-[4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-4-

Page 111 SAEED ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN methyl- (9CI) (CA INDEX NAME) (Continued)

Double bond geometry as shown.

RN 442561-55-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2)-[4-(2,6-difluorophenyl)-1,2-dihydro-2-oxo3H-indol-3-ylidene]methyl]-N-[2-(1,1-dioxido-4-thiomorpholinyl)ethyl]-4methyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continue No. 442561-76-4 CAPLUS COPYRIGHT 2005 ACS on STN (Continue No. 1H-Pyrrole-3-carboxamide, S-[(2)-(4-(4-chlorophenyl)-1,2-dhlydro-2-oxo-3H-indio-3-ylidene, methyl)-N-[3-(1,1-dioxido-4-thiomorpholinyl)-2-hydroxypropyl)-2,4-dimethyl- (9CI) (CA INDEX NAME) (Continued)

Double bond geometry as shown.

442561-77-5 CAPLUS
1H-Pyrrole-3-carboxamide,
[(2)-[4-(2,6-difluorophenyl)-1,2-dihydro-2-oxo3H-indol-3-ylidene]methyl]-N-(3-(1,1-dioxido-4-thiomorpholinyl)-2hydroxypropyl]-4-methyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 58 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 442561-56-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(2]-[4-(2,6-difluorophenyl)-1,2-dihydro-2-oxo3H-indol-3-ylidene]methyl]-N-[2-(1,1-dioxido-4-thiomorpholinyl)ethyl]-2fluoro-4-methyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 59 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2002:31440 CAPLUS
DOCUMENT NUMBER: 136:102386
4-heteroary1-3-heteroary1ideny12-indolinones and their use as protein kinase inhibitors
INVENTOR(S): Tang, Peng Cho; Wei, Chung Chen; Huang, Ping; Cui, Jingron
PATENT ASSIGNEE(S): Sugen, Inc., USA
SOURCE: PCT Int. Appl., 164 pp.
CODEN: PIXXD2
DOCUMENT TYPE: CODEN: PIXXD2
DATENT ASSIGNEE(S): Sigen, Inc., USA
FAMILY ACC. NUM. COUNT: 1
EARGUAGE: English
FAMILY ACC. NUM. COUNT: 1
FAMILY ACC. NUM. COUNT: 1

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

US 2001-894902 A3 20010629

WO 2001-US20768 W 20010629

OTHER SOURCE(S): MARPAT 136:102386

L4 ANSWER 59 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

AB Title compds. I (R1-2 = H, alkyl, cycloalkyl, aryl, heteroaryl,
heteroalicyclic, halo, etc.; Het = (un)substituted aromatic heterocycle
containing at least one and not more than two N atoms,
tetrahydro(thio)pyranyl, (thio)morpholino, piperidinyl, piperazinyl,
tetrazolyl, etc.: Q = (un)substituted aromatic heterocycle containing
not more
than two N atoms, 5-membered ring (un)substituted heterocycle containing
N, O

than two N atoms, 5-membered ring (un)substituted heterocycle containing N, O

or S, e.g., imidazolyl, pyrrolyl, indolyl, etc.) with some exceptions, were prepared Included are 75 synthetic examples and results for several protein tyrosine kinase assays for those compels. For instance, 4-bromoindole was coupled to bis(pinacolato)diborane (DMSO, KOAC, PdCl2(dppf)=KERCL2, 80°C, 22 hl. The resulting dioxaborolane was coupled to 4-bromopyridine=HCl (THF, Pd(PFN3)4, NAOH, 70°C, 6 h) to give the indole which was treated with CSHSN=873 (t-BuOH/ECOH/HZO, lh) followed by zinc (stirred 1 addnl. hour) to give 4-(pyridin-4-y1)-1,3-dyhydroindol-2-one as a yellow solid. Condensation of this intermediate with 5-methylimidazole-4-carboxaldehyde (ELON, piperidine, 2 days) afforded II. If had ICSO = 4.88 mM for FGFR-1 tyrosine kinase and 0.03 mM for cdk2/cyclin A tyrosine kinase. I are useful in treating cancer, immunol. disorders, etc.

II 388116-64-12 388117-08-07 388117-10-0P
388117-12-22 388117-08-07 388117-10-0P
388117-12-29 388117-08-07 388117-10-0P
388117-12-29 388117-08-07 388117-10-0P
(Uses)
(Uses)
(drug; preparation and use of 4-heteroaryl-3-heteroaryl-idenyl-2-indolinones and their use as protein kinase inhibitors)

RN 38816-64-1 CAPIUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(dicthylaminolethyl]-5-[[1,2-dihydro-2-oxo-4-(4-pyridinyl)-3H-indol-3-ylidene]methyl]-2, 4-dimethyl- (9CI) (CA INDEX NAME)

RN 388116-68-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[1,2-dihydro-2-oxo-4-(4-pyridinyl)-3H-indol-3ylidene]methyl]-2-methyl-4-phenyl-M-4-pyridinyl- (9CI) (CA INDEX NAME)

L4 ANSWER 59 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

388117-08-6 CAPLUS
1H-Pyrrole-3-carboxamide,
[1,2-dihydro-2-oxo-4-(4-piperidinyl)-3H-indol3-ylidene]methyl)-4-methyl-N-(phenylmethyl)- (9CI) (CA INDEX NAME)

388117-10-0 CAPLUS
1H-Pyrrole-3-carboxamide,
[1,2-dihydro-2-oxo-4-(4-piperidinyl)-3H-indol3-ylidene|methyl]-4-methyl-N-(4-pyridinylmethyl)- (9CI) (CA INDEX NAME)

L4 ANSWER 59 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 388116-99-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino|ethyl]-5-[{1,2-dihydro-2-oxo-4-(4-piperidinyl)-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 388117-02-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[1,2-dihydro-2-oxo-4-(4-piperidiny1)-3H-indol3-ylidene}methyl}-2-methyl-4-phenyl-N-4-pyridinyl- (9CI) (CA INDEX NAME)

L4 ANSWER 59 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 388117-12-2 CAPLUS
RN 188117-12-2 CAPLUS
RN

RN 388117-28-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(4-(2-amino-5-pyrimidinyl)-1,2-dihydro-2-oxo3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI)
(CA INDEX NAME)

388117-30-4 CAPLUS
1H-Pyrrole-3-carboxamide,
2-(diethylamino)ethyl)-5-[[1,2-dihydro-2-oxo4-(3-pyridinyl)-3H-indol-3-ylidene]methyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

TITLE: Mannich base prodrugs of 3-(pyrrol-2-ylmethylidene)-2INVENTOR(S): indolinone derivatives
MOON, Malcolm Wilson; Morozowich, Walter; Gao, Ping;
Tang, Peng Cho
Sugen, Inc., USA; Pharmacia & Upjohn Company
SOURCE: PCT Int. Appl., 96 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: PAHILY ACC. NUM. COUNT: 3 DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. PATENT NO.

WO 2001090068
WI AE, AG, AL,
CO. CR. CU,
GM, HR, HU,
LS, LT, LU,
RO, RU, SD,
UZ, VN, YU,
RW: GH, GM, KE,
BJ, CF, CG,
AU301064885
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EP 1301505
R: AT, BE, CH,
R: AT, BE, CH, KIND DATE APPLICATION NO. DATE AZ 20011129 VA
A3 20020606
AM, AT, AU, AZ, BA,
CZ, DE, DK, DM, DZ,
LV, MA, MD, MG, MK,
SE, SG, SI, SK, SI,
ZA, ZW, AM, AZ, BY,
LS, MW, MZ, SD, SI,
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A, EC, EE, ES, FI, GB, GD, GE, GH,
A, KE, KG, KP, KR, KZ, LC, LK, LR,
A, MN, MW, MX, MZ, NO, NZ, PL,
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KG, KZ, MD, RU, TJ, TM
A, SZ, TZ, UG, ZW, AT, BE, CH, CY,
AT, TH, LU, MC, NL, PT, SE, TR, BF,
ML, MR, NE, SN, TD, TG
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R: AT, BE, CH, DE, DK, ES, FR,
IE, SI, LT, LV, FI, RO, MK,
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20020916 US 6716870 US 2004127542 US 2003-429895 US 2003-743909 US 2000-207000P 20030505 US 2004127544 20040701 20031224 P 20000524 PRIORITY APPLN. INFO.: US 2000-225045P P 20000811 US 2001-863804 A1 20010524 US 2001-863819 A3 20010524 US 2001-863905 A1 20010524

L4 ANSWER 60 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN ACCESSION NUMBER: 2001:868415 CAPLUS DOCUMENT NUMBER: 136:697

Mannich base prodrugs of

DOCUMENT NUMBER:

L4 ANSWER 60 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) WO 2001-US16757 W 20010524

US 2002-243663 B1 20020916

OTHER SOURCE(S): MARPAT 136:697

AB The present invention is directed to Mannich base prodrugs of certain 3-(pyrrol-2-ylmethylidene)-2-indolinone derivs. that modulate the

3-(pyrrol-2-ylmethylidene)-2-indolinone derivs. that modulate the activity of protein kinases ("PKs"). Pharmaceutical compns. comprising these compds., methods of treating diseases related to abnormal PK activity utilizing pharmaceutical compns. comprising these compds. and methods of preparing them are also disclosed.

IT 557795-19-49
RL: RCT (Reactant): SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (Mannich base prodrugs of 3-(pyrrol-2-ylmethylidene)-2-indolinone derivs.)
RN 557795-19-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

375798-55-3P
RL: SPN (Synthetic preparation); PREP (Preparation)
(Mannich base prodrugs of 3-(pyrrol-2-ylmethylidene)-2-indolinone
derivs.)
375798-55-3 CAPLUS
1H-Pyrrole-3-carboxamide, N-[3-(diethylamino)-2-hydroxypropyl]-5-[{2}-{5fluoro-1, 2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN ACCESSION NUMBER: 2001:617993 CAPLUS COLUMENT NUMBER: 135:195497
TITLE: Preparation of pyriole substitu 135:195497
Preparation of pyrrole substituted 2-indolinone protein kinase inhibitors for treatment of cancer Tang, Peng Cho; Miller, Todd; Li, Xiaoyuan; Sun, Li; Wei, Chung Chen; Shirazian, Shahrzad; Liang, Congxin; Vojkovsky, Tomas; Nematalla, Assad S. Sugen, Inc., USA
PCT Int. Appl., 225 pp.
CODEN: PIXXD2
Patent INVENTOR (S): PATENT ASSIGNEE (S): SOURCE: DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION: Patent English 1 PATENT NO. KIND DATE APPLICATION NO. DATE WO 2001060814 WO 2001060814 W: AE, AG A3, AM, DE, IN, MD, SM, LS, FI, CI, AA B2 A2, LV, T2 AA AAAA1 20010823 WO 2001-US4813 20010215 20020124 , AU, AZ, , DM, DZ, , JP, KE, , MK, MN, , SI, TJ, , BY, KG, , MZ, SD, , GB, GR, , GA, GN, 20010823 20021024 20030603 20021113 2001060814
W: AE, AG, AL,
CR, CU, CZ,
HU, ID, IL,
LU, LV, MA,
SD, SE, SG,
YU, ZA, ZW,
RW: GH, GM, KE,
DE, DK, ES,
BJ, CF, CG, AT, DK, IS, MG, SK, AZ, MW, FR, CM, CA 2399358 US 2002156292 US 6573293 EP 1255752 EP 2001-914376 20010215
GB, GR, IT, LI, LU, NL, SE, MC, PT,
CY, AL, TR
BR 2001-8394 20010215
NO 2002-3831 20020813
ZA 2002-6459 20020813
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R: AT, BE, CH,

IE, SI, LT,

JP 2003523340

BR 2001008394

NO 2002003831

ZA 2002006469

BG 107079 20021113 , ES, FR, , RO, MK, 20030805 20040622 20021015 20031113 DK, FI, US 2004063773 PRIORITY APPLN. INFO.:

OTHER SOURCE(S): MARPAT 135:195497

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A3 20010215

W 20010215

The title compds. (I) [wherein Rl = H, halo, (cyclo)alkyl, (hetero)aryl, heteroalicyclic, OH, alkoxy, acyl, (un)aubstituted amino or carbamoyl, etc.; R2 = H, halo, alkyl, trihalomethyl, OH, alkoxy, CN, (hetero)aryl, (un)substituted amino, acyl (amino), or sulfamoyl, etc.; R3 = H, halo, alkyl, trihalomethyl, OH, alkoxy, (hetero)aryl, (un)substituted acyl, (acyl)amino, sulfamoyl, or alkylsulfonyl, etc.; R4 = H, halo, alkyl, OH, alkoxy, or (un)substituted acyl, (acyl)amino, sulfamoyl, or alkylsulfonyl, etc.; R4 = H, halo, alkyl, OH, alkoxy, or (un)substituted amino; R5 and R6 = independently R, alkyl, or acyl; R7 = H, alkyl, (hetero)aryl, or acyl; and their pharmaceutically acceptable salts] were prepared as protein kinase modulators for the treatment of cellular disorders such as cancer. For example, 5-fluoro-1,3-dhlydroindol-2-one was condensed with 5-formyl-2,4-dimethyl-lH-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide to give II (551). II exhibited comparable activity against Flk-1 and PDOFRB and inhibited PDOF-dependent receptor phosphorylation in cells with an IC50 value of approx. O.03 µM. In efficacy expts. against various cancers in mice, II was well tolerated at 80 mg/kg/day, even when dosed continuously for more than 100 days.

557795-19-49

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); TRU (Therapeutic use); BIOL (Biological study); PREF (Preparation); TRU (Therapeutic use); BIOL (Biological study); Operaration of pyrrole substituted 2-indolinone protein kinase by condensation of dibydroindolones with formylpyrroles for treatment

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(preparation of pyrrole substituted 2-indolinone protein Almee' inhibitors

by condensation of dihydroindolones with formylpyrroles for treatment of cancer and other diseases)

RN 557795-19-4 CAPFUS

CN 1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX MAMPY)

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN 356069-74-4P 356069-75-5P 356069-76-6P 356069-77-7P (Continued)

326914-10-7 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(Z)-{5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-{2-{diethylamino}ethyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

326914-17-4 CAPLUS
1H-Pyrrole-3-carboxamide,
-(dimethylamino)ethyl]-5-[(2)-(5-fluoro-1,2dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX
NAME)

Double bond geometry as shown.

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) Double bond geometry as shown.

126914-09-4P 326914-10-7P 326914-17-4P 326914-19-6P 342641-50-3P 342641-51-6P 342641-49-0P 342641-55-3P 342641-55-6P 342641-55-6P 342641-55-6P 342641-55-6P 342641-55-6P 342641-55-6P 342641-55-6P 342641-65-P 342641-70-P 342641-70-P 342641-70-P 342641-70-P 342641-71-9P 342641-81-9P 342641-81-9P 342641-81-9P 342641-91-9P 342661-91-9P 342661-91-

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

326914-19-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(1,2-dihydro-2-oxo-3H-indol-3ylidene)methyl]-N-(2-(dimethylamino)ethyl)-2,4-dimethyl- (SCI) (CA INDEX NAME)

342641-49-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[3-(diethylamino)propyl]-2-(1-methylethyl)-4-phenyl-(SCI) (CA INDEX NAME)

342641-50-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2-(1-methylethyl)-4-phenyl-N-[3-(1-pyrrolidinyl)propyl]-(9CI) (CA INDEX NAME)

RN 342641-51-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-yliden)methyl]-N-[2-(diethylamino)ethyl]-2-(1-methylethyl)-4-phenyl-(9CI) (CA INDEX NAME)

RN 342641-52-5 CAPLUS RN lH-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2-(1-methylethyl)-N-[3-(4-methyl-1-piperazinyl)propyl]-4-phenyl- (9CI) (CA INDEX NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continue

RN 342641-56-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(dimethylamino)ethyl]-2-methyl-4-phenyl- (9CI) (CA INDEX NAME)

RN 342641-57-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{[1,2-dihydro-6-(2-methoxyphenyl)-2-oxo-3H-indol-3-ylidene|methyl]-N-[2-(dimethylamino)ethyl]-2-methyl-4-phenyl-(9CI) (CA INDEX NAME)

RN 342641-59-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-((5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-54-7 CAPLUS

(N IH-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2-methyl-4-phenyl-N-[2-(1-pyrrolidinyl)ethyl}- (9CI) (CA INDEX NAME)

RN 342641-55-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{[1,2-dihydro-6-(2-methoxyphenyl)-2-oxo-3H-indol-3-ylidene|methyl)-2-methyl-4-phenyl-N-[2-(1-pyrrolidinyl)ethyl)-(9CI) (CA INDEX NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene)methyl]-N-[3-(diethylamino)propyl]-2-methyl-4-phenyl- (9CI) (CA INDEX NAME)

RN 342641-60-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-61-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-yliden)methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-62-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

342641-63-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-64-9 CAPLUS
CN 1H-Fyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

342641-68-3 CAPLUS
1H-Pyrrole-3-carboxamide,
-(diethylamino)ethyl)-5-[(1,2-dihydro-2-oxo5-phenyl-3H-indol-3-ylidene)methyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

342641-69-4 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA

342641-70-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl]-N-(3-(1H-imidazol-1-yl)propyl]-2, 4-dimethyl- (9Cl) (CA

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-65-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[3-(1H-imidazol-1-yl)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

342641-66-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{1,2-dihydro-6-(2-methoxyphenyl)-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

342641-67-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-(3-methoxyphenyl)-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN INDEX NAME) (Continued)

342641-71-8 CAPLUS
1H-Pyrrole-3-carboxamide,
2-(diethylamino)ethyl)-5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-73-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-N-[3-(1H-imidazol-1-yl)propyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-74-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[6-[3,5-dichlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene]methyl]-N-[2-[diethylamino]ethyl]-2,4-dimethyl- (9CI)
(CA
INDEX NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued

RN 342641-77-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[{1,2-dihydro-2-oxo-6-(3-pyridinyl)-3H-indol-3-ylidene|methyl|-N-[3-(dimethylamino)propyl]-2,4-dimethyl- (9CI) (CA INDEX
NAME)

RN 342641-78-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl)-N-[3-(dimethylamino)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-75-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(dicthylamino|ethyl)-5-[[1,2-dihydro-2-oxo-6-(3-pyridinyl)-3H-indol-3-ylidene]methyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

RN 342641-76-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[{1,2-dihydro-2-oxo-6-(3-pyridinyl)-3H-indol-3-ylidene|methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX
NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 342641-79-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-{3-(ditchylaminol)propyl}-5-{(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl}-2,4-dimethyl- {9CI} (CA INDEX NAME)

Ph CH CH2) 3-NEt2

RN 342641-80-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[3-(dicthylamino)propyl]-5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Ph H O H Me
CH NH- (CH<sub>2</sub>) 3-NEt<sub>2</sub>

RN 342641-81-0 CAPLUS
CN 1H-Indole-4-carboxamide, N-(3-chloro-4-methoxyphenyl)-3-[[4-[[[3-(diethylamino)propyl)amino]carbonyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-(9CI) (CA INDEX NAME)

C1 OMe

RN 342641-82-1 CAPLUS CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

Page 118 SAEED

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene)methyl]-N-[3-(diethylamino)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-83-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-n-[2-(diethylamino)ethyl]-2,4-bis(1-methylethyl)- (9CI) (CA INDEX NAME)

RN 342641-84-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[3-(diethylamino)propyl]-2,4-bis(1-methylethyl)- (9CI) (CA INDEX NAME)

RN 342641-85-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-bis(1-methylethyl)-N-[3-(1-pyrrolidinyl)propyl](9CI)
(CA INDEX NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continu

RN 342641-89-8 CAPLUS CN 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-[2-methoxy-5-[1-

methylethyl)phenyl}-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

RN 342641-91-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[6-(4-ethylphenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene|methyl|-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-87-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl-N-(4-pyridinylmethyl)- (9CI) (CA INDEX NAME)

RN 342641-88-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[6-(4-buty]phenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene|methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-92-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[6-{2, 4-dimethoxyphenyl}-1, 2-dihydro-2-oxo-3H-indol-3-ylidene|methyl}-2, 4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

RN 342641-93-4 CAPLUS
CN lH-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-[3-(1-methylethyl)phenyl]-2-oxo-3M-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]-(9CI) (CA INDEX NAME)

342641-94-5 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-95-6 CAPLUS
CN 1H-Indole-6-carboxylic acid,
3-[4-[[2-(diethylamino)ethyl]amino]carbonyl
|-3,5-dimethyl-lH-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo- [9CI] (CA
INDEX NAME)

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) pyridinylamino|sulfonyl|-3H-indol-3-ylidene|methyl|-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

342642-01-7 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[[5-

[(dimethylamino)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

342642-02-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[5-[([3-chlorophenyl)amino]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-96-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[5-[(dimethylamino)sulfonyl]-1,2-dihydro-2-oxo-3H-indd)-3-ylidene]methyl]-2,4-dimethyl-N-[2-(l-pyrrolidinyl)ethyl]-(9CI) (CA INDEX NAME)

342641-97-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[5-[[(3-chlorophenyl)amino]sulfonyl]-1,2-dhydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-[1-pyrrolidinyl]ethyl]- (9C1) (CA INDEX NAME)

342641-98-9 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[{1,2-dihydro-2-oxo-5-[(3-

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 342642-09-5 CAPLUS Glycine, N-[(5-[4],2-dihydro-4-methyl-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-yildene|methyl]-4-methyl-1H-pyrrol-3-yilcarbonyl]-, ethyl ester (9CI) (CA INDEX NAME)

342642-10-8 CAPLUS Glycine, N-[{5-[[1,2-dihydro-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-ylidene]methyl]-4-methyl-1H-pyrrol-3-yl)carbonyl}-, ethyl ester (9CI)

INDEX NAME)

342642-11-9 CAPLUS
Glycine, N-[[5-[[1,2-dihydro-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3ylidene]methyl]-4-methyl-1H-pyrrol-3-yl]carbonyl]- (9CI) (CA INDEX NAME)

346405-32-1 CAPLUS

1H-Pyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-yildene)methyll-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA

INDEX

RN 356068-82-1 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAMES)

356068-90-1 CAPLUS 1H-Pytrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 356068-94-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(2]-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA
INDEX

Double bond geometry as shown.

356068-95-6 CAPLUS
1H-Pyrrole-3-carboxamide,
(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]N-{2-(dimethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

356068-96-7 CAPLUS
lH-Pyrrole-3-carboxamide, N-[2-(dimethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-(9CI) (CA INDEX

356068-97-8 CAPLUS

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 356068-91-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino) ethyl]-5-[(1,2-dihydro-2-oxo3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 356068-92-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
1H-Pyrrole-3-carboxamide, N-[2-(ethylamino)ethyl]-5-([2]-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 356068-99-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-[diethyloxidoamino]ethyl]-5-[(Z)-(5-fluoro1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-03-9 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-[2-(4-methyl-1-piperazinyl)ethyl}- (9CI)(CA INDEX NAME)

356069-04-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(4-methyl-1-piperarinyl)ethyl]- (9CI)(CA INDEX NAME)

356069-05-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN L4

356069-09-5 CAPLUS'
1H-Pyrrole-3-carboxamide, 5-{(Z)-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-[2-(3,5-dimethyl-1-piperazinyl)ethyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

 $356069-12-0 \quad CAPLUS \\ 1H-Pytrole-3-carboxamide, \quad N-\{2-\{3,5-dimethyl-1-piperazinyl\}ethyl\}-5-[\{2\}-\{5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1-2$ 

(9CI)

(CA INDEX NAME)

Double bond geometry as shown.

356069-13-1 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(3,5-dimethyl-1-piperazinyl)ethyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene)methyl]-2,4-dimethyl-N-[2-(4-methyl-1-piperazinyl)ethyl]- (9CI) (CA INDEX NAME)

RN 356069-07-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]2,4-dimethyl-N-[2-(4-methyl-1-piperazinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN Double bond geometry as shown.

356069-15-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(3,5-dimethyl-1-piperazinyl)ethyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 356069-16-4 CAPLUS
CN 1R-Pyrcole-3-carboxamide,
N-[2-(ethylmaino) ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- [9CI] (CA INDEX NAME)

RN 356069-17-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethyloxidoamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 356069-18-6 CAPLUS
CN Butanedioic acid, hydroxy-, (2S)-, compd. with 5-[{5-chloro-1,2-dihydro-2-

oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl}-1H-pyrrole-3-carboxamide (9CI) (CA INDEX NAME)

CM 1

CRN 356068-82-1 CMF C22 H25 C1 N4 O2

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

CH2-CH2-NEt2

CM 2

Absolute stereochemistry. Rotation (-).

HO<sub>2</sub>C

RN 356069-20-0 CAPLUS
CN Butanedioic acid, hydroxy-, (2S)-, compd. with
5-[(5-chloro-1,2-dihydro-2oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl-1Hpyrrole-3-carboxamide (9CI) (CA INDEX NAME)

CM 1

CRN 356068-90-1 CMF C22 H27 C1 N4 O2

-CH2-CH2-NEt2

CM 2

Absolute stereochemistry. Rotation (-).

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

Absolute stereochemistry. Rotation (-).

RN 356069-19-7 CAPLUS
CN Butanedioic acid, hydroxy-, {2S}-, compd. with
N-{2-(diethylamino)ethyl}-5[(5-fluoro-1,2-dhydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-1Hpyrrole-3-carboxamide (9CI) (CA INDEX NAME)

CM 1

CRN 342641-94-5 CMF C22 H27 F N4 O2

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 356069-21-1 CAPLUS
CN Butanedioic acid, hydroxy-, (2S)-, compd. with
5-{(5-fluoro-1,2-dihydro-2-

oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]-lH-pyrrole-3-carboxamide (9CI) (CA INDEX NAME)

CM 1

CRN 346405-32-1 CMF C22 H25 F N4 O2

CM 2

Absolute stereochemistry. Rotation (-).

CO2H но₂с′

356069-22-2 CAPLUS
Butanedioic acid, hydroxy-, (2s)-, compd. with N-[2-(ethylamino)ethyl]-5[(5-fluoro-1,2-dihydro-2-oxo-3M-indol-3-ylidene)methyl)-2,4-dimethyl-1Hpyrrole-3-carboxamide (9CI) (CA INDEX NAME)

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SAEED

2

Absolute stereochemistry. Rotation (-).

356069-23-3 CAPLUS
Butanedioic acid, hydroxy-, (2S)-, compd. with N-[2-(diethyloxidoamino)ethyl]-5-[(2]-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrole-3-carboxamide (9CI) (CA INDEX NAME)

CM 1

CRN 356068-99-0 CMF C22 H27 F N4 O3

Double bond geometry as shown.

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356069-27-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl-N-[3-(tetrahydro-2-oxo-1(2H)-pyrimidinyl)propyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-28-8 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[3-(tetrahydro-2-oxo-1(2H)-pyrimidinyl)propyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-29-9 CAPLUS lH-Pyrrole-3-carboxamide,  $5-\{(z)-\{5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene]$  methyl]-2, 4-dimethyl-N- $\{3-(tetrahydro-2-oxo-1(2H)-yrimidinyl)$  propyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

Absolute stereochemistry. Rotation (-).

356069-24-4 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(acetylamino)ethyl]-5-{(2)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-25-5 CAPLUS | H-Pyrrole-3-carboxamide, N-[2-(acetylamino)ethyl]-5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

Double bond geometry as shown.

356069-26-6 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(acetylamino)ethyl]-5-[(2)-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (SCI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356069-30-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{Z}]-{1,2-dihydro-2-oxo-3H-indol-3-ylidene| methyl]-2,4-dimethyl-N-[3-(tetrahydro-2-oxo-1{2H})-pyrimidinyl)propyl]- {9CI} (CA INDEX NAME)

Double bond geometry as shown.

356069-31-3 CAPLUS

HH-Pyrrole-3-carboxamide, 5-{(2)-(5-cyano-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-(3-(tetrahydro-2-oxo-1(2H)-pyrimidinyl)propyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-33-5 CAPLUS lH-Pyrrole-3-carboxamide,  $5-\{(Z)-\{5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene) methyl]-2,4-dimethyl-N-\{2-\{3-oxo-1-piperazinyl\}ethyl\}-, mono(trifluoroacetate) (9C1) (CA INDEX NAME)$ 

CM 1

CRN 356069-32-4 CMF C22 H24 Br N5 O3

CM 2

RN 356069-34-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{(Z)-{1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-{2-(1-pyrrolidinyl)ethyl}- {9CI} (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356069-38-0 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(2)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(2-oxo-1-imidazolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-39-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl-N-[2-(2-oxo-1-imidazolidinyl)ethyl]- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 356069-35-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX

Double bond geometry as shown.

356069-36-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-cyano-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[3-(2-oxo-1-pyrrolidinyl)propyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-37-9 CAPLUS
1H-Pytrole-3-carboxamide, 5-[(2)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(2-oxo-1-imidazolidinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

356069-40-4 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{{Z}}-{1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-{2-{2-oxo-1-imidazolidinyl}ethyl}- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-41-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(z)-(5-cyano-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(2-oxo-1-imidazolidinyl)ethyl]- (9CI)
(CA INDEX NAME)

356069-42-6 CAPLUS
1-Piperazineacetic acid, 4-[2-[[[5-[(Z)-(5-bromo-1,2-dihydro-2-oxo-3Hindol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]amino]ethyl], ethyl ester (SCI) (CA INDEX NAME)

Double bond geometry as shown.

356069-43-7 CAPLUS
1-Piperazineacetic acid, 4-[2-[[5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3Hindol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]amino]ethyl}, ethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356069-46-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(Z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3ylidene)methyl]-N-[3-(hexahydro-2-oxo-1H-azepin-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

356069-47-1 CAPLUS 1H-Pyrrole-3-carboxamide, 5-{(2)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3ylidene)methyl]-N-[3-(hexahydro-2-oxo-1H-azepin-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

356069-48-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3ylidene)methyl)-N-[3-(hexahydro-2-oxo-1H-azepin-1-yl)propyl)-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

356069-44-8 CAPLUS
1-Piperazineacetic acid, 4-[2-[[[5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-

indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]amino]ethyl], ethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-45-9 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-{(cyanomethyl)amino]ethyl}-5-[(Z)-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

$$\overline{\phantom{a}}$$

356069-49-3 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[{Z}]-{1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl)-N-[3-(hexahydro-2-oxo-1H-azepin-1-yl)propyl]-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown

356069-50-6 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-cyano-1,2-dihydro-2-oxo-3H-indol-3-

ylidene)methyl)-N-{3-{hexahydro-2-oxo-1H-azepin-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-51-7 CAPLUS
1H-Fyrrole-3-carboxamide, N-[2-(acetylamino)ethyl]-5-[(Z)-(5-bromo-1,2-dlhydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

356069-53-9 CAPLUS

1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-ylidene)methyl)-2,4-dimethyl-N-[2-(3-oxo-1-piperazinyl)ethyl)-,
mono(trifluoroacetate) (9C1) (CA INDEX NAME)

CM 1

CRN 356069-52-8 CMF C22 H24 F N5 O3

Double bond geometry as shown.

CM 2

CRN 76-05-1 CMF C2 H F3 O2

356069-55-1 CAPLUS

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

CM 2

356069-58-4 CAPLUS

1H-Pyrrole-3-carboxamide, 5-[(2)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3ylidene] methyl]-N-[2-[(2-cyanoethyl)amino]ethyl]-2,4-dimethyl- (9CI) (CA
INDEX NAME)

Double bond geometry as shown.

356069-59-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-[(2-cyanoethyl)amino]ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
CN 1H-Pyrrole-3-carboxamide, 5-{{2} - {1} - {1}, 2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2, 4-dimethyl-N-{2-{3}-oxo-1-piperazinyl}ethyl}-,
mono(trifluoroacetate) {9CI} (CA INDEX NAME)

CM 1

CRN 356069-54-0 CMF C22 H25 N5 O3

Double bond geometry as shown.

CM 2

356069-57-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-cyano-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(3-oxo-1-piperazinyl)ethyl]-,
mono(trifluoroacetate) (SCI) (CA INDEX NAME)

CM 1

CRN 356069-56-2 CMF C23 H24 N6 O3

Double bond geometry as shown.

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

356069-60-8 CAPLUS 1H-Pyrrole-3-carboxamide, N-[2-[(2-cyanoethyl)amino)ethyl]-5-{(Z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-61-9 CAPLUS IH-Pyrrole-3-carboxamide, N-[2-[(2-cyanoethyl)amino]ethyl]-5-[(2)-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX

Double bond geometry as shown.

356069-62-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-cyano-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-(2-[(2-cyanoethyl)amino]ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

356069-64-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(2)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylideno|methyl|-2,4-dimethyl-N-[2-(3-oxo-1-piperazinyl)ethyl]-,
mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CRN 356069-63-1 CMF C22 H24 C1 N5 03

Double bond geometry as shown.

2 CM CRN 76-05-1 CMF C2 H F3 O2

356069-65-3 CAPLUS

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

356069-68-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(z)-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(4-methyl-1-piperazinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-69-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{{Z}}-{1,2-dihydro-2-oxo-3H-indol-3-ylidene}methyl]-2,4-dimethyl-N-{3-(4-methyl-1-piperazinyl)propyl}- (9CI)
(CA INDEX NAME)

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
CN 1H-Pyrrole-3-carboxamide, 5-[(2)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3ylidene)methyl]-2,4-dimethyl-N-[2-(4-methyl-1-piperazinyl)ethyl]- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

356069-66-4 CAPLUS
1H-Pyrole-3-carboxamide, 5-{{Z}}-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-{2-(4-methyl-1-piperazinyl)ethyl}- (9CI)
(CA INDEX NAME)

Double bond geometry as shown.

356069-67-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl-N-[2-(4-methyl-1-piperazinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 356069-70-0 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(z)-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[3-(4-methyl-1-piperazinyl)propyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-71-1 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(2)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[3-(4-methyl-1-piperazinyl)propyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-72-2 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[(z)-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[3-(4-methyl-1-piperazinyl)propyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-73-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-{1,2-dihydro-2-oxo-3H-indol-3-yiidene|methyl]-2,4-dimethyl-N-[2-[4-(phenylmethyl)-1-piperazinyl]ethyl]-(9CI) (CA INDEX NAME)

356069-74-4 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{Z}]-(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-[4-(phenylmethyl)-1-piperazinyl]ethyl)-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-75-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(Z)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl)-2,4-dimethyl-N-[2-[4-(phenylmethyl)-1-piperazinyl}ethyl]-(GCI (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

L4 ANSWER 61 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

356069-76-6 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[{2}-(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-[4-(phenylmethyl)-1-piperazinyl]ethyl]-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

356069-77-7 CAPLUS
1-Piperazineacetic acid, 4-[2-{[[5-[{Z}]-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]amino]ethyl]-, ethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4 ANSWER 62 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2001:472477 CAPLUS

DOCUMENT NUMBER: 135:56059 Methods of modulating c-kit tyrosine protein kinase function with indolinone compounds

INVENTOR(S): Lipson, Ken; McMahon, Gerald

SUBJOURNITY ASSIGNEE(S): Sugen, Inc., USA

POT Int. Appl., 59 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

|      |     |      |      |      |     |     |     |      |      |     |      | LICAT          |      |     |     |      |       |     |
|------|-----|------|------|------|-----|-----|-----|------|------|-----|------|----------------|------|-----|-----|------|-------|-----|
|      |     |      |      |      |     |     |     |      |      |     |      | 2000-          |      |     |     |      |       |     |
|      |     | 2001 |      |      |     |     |     |      |      |     |      | -000-          | 0333 | 003 |     | -    | .0001 | 222 |
|      |     |      |      |      |     |     |     |      |      |     |      |                |      |     |     | ~    |       |     |
|      |     | ₩:   |      |      |     |     |     |      |      |     |      | BG,            |      |     |     |      |       |     |
|      |     |      |      |      |     |     |     |      |      |     |      | FI,            |      |     |     |      |       |     |
|      |     |      |      |      |     |     |     |      |      |     |      | KR,            |      |     |     |      |       |     |
|      |     |      |      |      |     |     |     |      |      |     |      | M2,            |      |     |     |      |       |     |
|      |     |      |      |      |     |     |     |      |      |     |      | TT,            |      |     | υG, | US,  | UZ,   | vn, |
|      |     |      |      |      |     |     |     |      |      |     |      | RU,            |      |     |     |      |       |     |
|      |     | RW:  |      |      |     |     |     |      |      |     |      | TZ,            |      |     |     |      |       |     |
|      |     |      |      |      |     |     |     |      |      |     |      | LU,            |      |     |     |      |       | BF, |
|      |     |      |      |      |     |     |     |      |      |     |      | MR,            |      |     |     |      |       |     |
|      |     |      |      |      |     |     |     |      |      |     |      | -000           |      |     |     |      |       |     |
|      |     |      |      |      |     |     |     |      |      |     |      | -000           |      |     |     |      |       |     |
|      | EΡ  | 1255 | 536  |      |     | A2  |     | 2002 | 1113 | 1   | EP 2 | -000           | 9917 | 04  |     | 2    | 0001  | 222 |
|      |     | R:   | AT,  | Β£,  | CH, | DE, | DK, | ES,  | FR,  | GB, | GR,  | IT,            | LI,  | LU, | NL, | SE,  | MC,   | PT, |
|      |     |      | IE,  | SI,  | LT, | LV, | FI, | RO,  | MK,  | CY, | AL,  | TR             |      |     |     |      |       |     |
|      | JΡ  | 2004 | 5003 | 63   |     | T2  |     | 2004 | 0108 |     | JP 2 | 2001-          | 5464 | 28  |     | 2    | 0001  | 222 |
|      | ΝZ  | 5196 | 97   |      |     | A   |     | 2004 | 0827 | 1   | NZ 2 | 2000-<br>2003- | 5196 | 97  |     | 2    | 0001  | 222 |
|      | US  | 2004 | 0025 | 34   |     | A1  |     | 2004 | 0101 | 1   | us 2 | 2003-          | 6008 | 68  |     | 2    | 0030  | 623 |
| PRIO | RIT | APP  | LN.  | INFO | .:  |     |     |      |      | 1   | US : | 1999-          | 1716 | 93P | 1   | P 1  | 9991  | 222 |
|      |     |      |      |      |     |     |     |      |      |     |      | 2000-          | 7410 | 42  |     |      | 0001  | 222 |
|      |     |      |      |      |     |     |     |      |      |     | 03 4 | -000           | ,410 | 4.6 |     | 31 Z | 10001 | 222 |
|      |     |      |      |      |     |     |     |      |      | 1   | NO 2 | 2000-          | US35 | 009 | 1   | 4 2  | 0001  | 222 |

OTHER SOURCE(S): MARPAT 135:56059

AB The invention concerns indolinone compds. and their use to inhibit the activity of a receptor tyrosine kinase. The invention is preferably used to treat cell proliferative disorders such as cancers characterized by over-activity or inappropriate activity of c-kit kinase.

IT 34264-94-5 34640-32-1

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES

USES

(Uses)
(indolinone derivs. for c-kit tyrosine protein kinase function modulation)
342641-94-5 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-{(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 346405-32-1 CAPLUS
CN 1H-Fyrrole-3-carboxamide, 5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA

PATENT NO. KIND DATE APPLICATION NO. DATE WO 2001037820 AZ 20010531 WO 2000-US32277 20001122

W1 AE, AG, AL, AM, AT, AU, AZ, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MK, MZ, NC, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VM, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CH, GA, GN, GW, ML, MR, NE, SN, TD, TG

EP 1233943 AZ 20020828 EP 2000-982228 C0001122

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LIT, LU, NL, SE, MC, PT, JF 2003514851 T2 20030422 JF 2001-539435 20001122

PRIORITY APPLN: INFO: WO 2000-US32277 W 20001122 MARPAT 135:19549

135:19549
Preparation of pyrrole substituted 2-indolinones as antitumor agents
Santitumor agents
Sugen, Inc., USA
PCT Int. Appl., 249 pp.
CODEN: PIXXD2
Patent

L4 ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS ON STN ACCESSION NUMBER: 2001:396655 CAPLUS DOCUMENT NUMBER: 135:19549

English

DOCUMENT NUMBER:

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

TITLE: INVENTOR (S): PATENT ASSIGNEE(S): SOURCE: DOCUMENT TYPE:

OTHER SOURCE(S):

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(CH<sub>2</sub>)<sub>3</sub>-NMe<sub>2</sub>

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
The title compds. (I: R1 = H, alkyl, alkenyl, etc.; R2 = H, halo, alkyl,
etc.; R3-R6 = H, alkyl, trihaloalkyl, etc.; R3 and R4, R4 and R5, R5 and
R6 may combine to form a six membered azyl ring, OCHZO, OCHZCHZO: R7 = H,
alkyl, cycloalkyl, etc.; R8-R1O = H, alkyl, trihaloalkyl, etc.] were
prepared and formulated. E.g., a multi-step synthesis of I [R1-R7 = H;

R10 = Me; R9 = (CH2)2CO2H) which showed 79-86% inhibition of tumor growth of Calu-6 cells in mice at 75 and 100 mg/kg/day, was given. The present invention features formulations of indolinones which compds. are

inventor teacher transfer and a street and a street action as free acids or free bases. The formulation is suitable for parenteral or oral administration, wherein the formulation comprises an ionizable substituted indolinone, and a pharmaceutically acceptable carrier therefor. The term "ionizable substituted indolinone" includes pyrrole substituted 2-indolinones I which, in addition to being otherwise

optionally substituted on both the pyrrole and 2-indolinone portions of the

compound,
are necessarily substituted on the pyrrole moiety with one or more
hydrocarbon chains which themselves are substituted with at least one

are necessarily substituted on the pyrrole moiety with one or more hydrocarbon chains which themselves are substituted with at least one polar group are substituted are substituted with at least one polar group are substituted are substit

(CH<sub>2</sub>)<sub>3</sub>-NMe<sub>2</sub>

251356-65-7 CAPLUS
2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-6-(2-methoxyphenyl)- (9CI) (CA INDEX NAME)

251356-63-5 CAPLUS 2H-Indol-2-one, 5-bromo-3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl|methylene|-1,3-dihydro- (9CI) (CA INDEX NAME)

251356-64-6 CAPLUS
2H-Indol-2-one, 3-[[4-{3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene}-1,3-dihydro-6-phenyl- (9CI) (CA INDEX NAME)

251356-66-8 CAPLUS
2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-6-(3-methoxyphenyl)- (9CI) (CA INDEX NAME)

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(Continued)

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

251356-67-9 CAPLUS 2H-Indol-2-one. 3-{[4-{3-(dimethylamino)propyl}-3,5-dimethyl-1H-pyrrol-2-yllmethylene]-1,3-dihydro-6-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)

251356-68-0 CAPLUS

2H-Indol-2-one, 5-chloro-3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro- (9CI) (CA INDEX NAME)

251356-69-1 CAPLUS
2H-Indol-2-one, 6-chloro-3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-lHpyrrol-2-yllmethylene]-1,3-dihydro- (9CI) (CA INDEX NAME)

251356-70-4 CAPLUS 2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 2H-Indol-2-one, 3-[(4-(3-(dimethylamino)propy)]-3,5-dimethyl-1H-pyrrol-2-yllmethylenej-1,3-dihydro-4-(2-hydroxyethyl)- (9CI) (CA INDEX NAME)

251356-75-9 CAPLUS
1H-Indole-5-sulfonamide, 3-([4-[3-(dimethylamino)propyl]-3,5-dimethyl-lHpyrrol-2-yl|methylene|-2,3-dihydro-2-oxo- (9CI) (CA INDEX NAME)

251336-76-0 CAPLUS
1H-Indole-5-sulfonamide, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-N-(1-methylethyl)-2-oxo- (9CI) (CA INDEX NAME)

251356-77-1 CAPLUS
MOTPholine, 4-[[3-[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-1H-indol-5-yl]sulfonyl]- (9CI) (CA INDEX NAME)

251356-78-2 CAPLUS 1H-Indole-5-sulfonamide, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) yl]methylene]-1,3-dihydro-5-methoxy- (9CI) (CA INDEX NAME)

251356-71-5 CAPLUS
2H-Indol-Z-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2yl]methylenej-1,3-dihydro-6-methoxy- (9CI) (CA INDEX NAME)

251356-72-6 CAPLUS
2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2yl]methylenej-1,3-dihydro-5-methyl- (9CI) (CA INDEX NAME)

251356-73-7 CAPLUS 2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-4-methyl- (9CI) (CA INDEX NAME)

RN 251356-74-8 CAPLUS

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
Pyrrol-2-yl]methylenej-2,3-dihydro-N,N-dimethyl-2-oxo- (9CI) (CA INDEX NAME)

342641-49-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[3-(diethylamino)propyl]-2-(1-methylethyl)-4-phenyl-(9CI) (CA INDEX NAME)

342641-50-3 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2-(1-methylethyl)-4-phenyl-N-(3-(1-pyrrolidinyl)propyl]-(9CI) (CA INDEX NAME)

342641-51-4 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2-(1-methylethyl)-4-phenyl-(9CI) (CA INDEX NAME)

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342641-52-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2-(1-methylethyl)-N-(3-(4-methyl-1-piperazinyl)propyl}-4-phenyl- (9CI) (CA INDEX NAME)

342641-54-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2-methyl-4-phenyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-57-0 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-(2-methoxypheny1)-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(dimethylamino)ethyl]-2-methyl-4-phenyl-(9CI) (CA INDEX NAME)

342641-59-2 CAPLUS IH-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[3-(diethylamino)propyl]-2-methyl-4-phenyl- (9CI) (CA INDEX NAME)

342641-60-5 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-{2-(dimethylamino)ethyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-55-8 CAPLUS
1H-Pyrrole-3-carboxemide, 5-[[1,2-dihydro-6-(2-methoxypheny1)-2-oxo-3H-indol-3-ylidene]methyl]-2-methyl-4-phenyl-N-[2-(1-pyrrolidinyl)ethyl]-(9CI) (CA INDEX NAME)

342641-56-9 CAPLUS
1H-Pytrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-{2-(dimethylamino)ethyl}-2-methyl-4-phenyl- (9CI) (CA INDEX NAME)

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-61-6 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl}-N-{2-(dimethylamino)ethyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

342641-62-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-[2-(dimethylamino)ethyl}-2,4-dimethyl- (9CI) (CA INDEX

342641-63-8 CAPLUS IH-Pytrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

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RN 342641-64-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX

342641-65-0 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-N-{3-{lH-imidazol-1-yl}propyl}-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
CN 1H-Pyrrole-3-carboxamide,
N-[2-(dicthylamino)=thyl)-5-(1,2-dihydro-2-oxo5-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-69-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-{(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-{2-(1-pyrrolidinyl)ethyl}- (9CI) (CA

342641-70-7 CAPLUS
1H-Pyrrole-3-carboxamide, 5-{{1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-yylidene|methyl}-N-{3-(1H-imidazol-1-yl)propyl}-2,4-dimethyl-(9CI) (CA INDEX NAME)

L4 ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-66-1 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-(2-methoxypheny1)-2-oxo-3H-indol-3-ylidene]methyl]-N-[2-(dimethylamino)ethyl]-2,4-dimethyl- (SCI) (CA INDEX NAME)

342641-67-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-(3-methoxyphenyl)-2-oxo-3H-indol-3-ylidene|methyl}-N-[2-(dimethylamino)ethyl}-2,4-dimethyl- (9CI)
(CA INDEX NAME)

342641-68-3 CAPLUS

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-71-8 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[2-(diethylamino)=thyl]-5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-72-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX

RN 342641-73-0 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-((1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl)-N-(3-(1H-imidazol-1-yl)propyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-74-1 CAPLUS
CN 1H-Pytrole-3-carboxamide,
5-[[6-(3,5-dichlorophenyl)-1,2-dihydro-2-oxo-3Hindol-3-ylidene]methyl]-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI)
(CA
INDEX NAME)

L4 ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-77-4 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[1,2-dihydro-2-oxo-6-[3-pyridinyl)-3H-indol-3yliden]methyl]-N-[3-(dimethylamino)propyl]-2,4-dimethyl- (9CI) (CA
INDEX
NAME)

RN 342641-78-5 CAPLUS
CN 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-2-oxo-5-phenyl-3H-indol-3-ylidene)methyl]-N-[3-(dimethylamino)propyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-75-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-(2-(diethylamino)ethyl)-5-[[1,2-dihydro-2-oxo-6-(3-pyridinyl)-3H-indol-3-ylidene}methyl]-2,4-dimethyl- {9CI} (CA INDEX NAME)

RN 342641-76-3 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[{1,2-dh}ydro-2-oxo-6-(3-pyridiny1)-3H-indol-3ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrolidiny1)ethyl]- (9CI) (CA
INDEX
NAMP)

L4 ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
RN 342641-79-6 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[3-(dicthylaminol) propyl]-5-[(1,2-dihydro-2-oxo5-phenyl-3H-indol-3-ylidene) methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-80-9 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
N-[3-(dichtylamino)propy]]-5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-81-0 CAPLUS
CN 1H-Indole-4-carboxamide, N-(3-chloro-4-methoxyphenyl)-3-[[4-[[3-(diethylamino)propyl]amino]carbonyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-(9CI) (CA INDEX NAME)

RN 342641-82-1 CAPLUS CN 1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-

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ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) ylidene|methyl|-N-[3-(diethylamino)propyl|-2,4-dimethyl- (SCI) (CA INDEX NAME)

342641-83-2 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-2,4-bis(1-methylethyl)- (9CI)(CA INDEX NAME)

342641-84-3 CAPLUS 1H-Pyrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylideno|methyl]-N-[3-(diethylamino)propyl]-2,4-bis(1-methylethyl)- (9CI) (CA INDEX NAME)

RN 342641-85-4 CAPLUS
CN H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-bis(1-methylethyl)-N-[3-(1-pyrrolidinyl)propyl]-(9CI) (CA INDEX NAME)

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

342641-89-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-[2-methoxy-5-[1-

methylethyl)phenyl)-2-oxo-3H-indol-3-ylidene|methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

RN 342641-91-2 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[{6-(4-ethylphenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-87-6 CAPLUS

1H-Pyrrole-3-carboxamide, 5-{(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl}-2,4-dimethyl-N-(4-pyridinylmethyl)- {9CI} {CA INDEX

RN 342641-88-7 CAPLUS
CN 1H-Pyrrole-3-carboxamide,
5-[[6-(4-butylphenyl)-1,2-dihydro-2-oxo-3H-indol3-ylidene|methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

RN 342641-92-3 CAPLUS
CN 1H-Pytrole-3-carboxamide,
[{6-(2,4-dimethoxyphenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene|methyl]-2,4-dimethyl-N-{2-(1-pytrolidinyl)ethyl]- (9CI)
(CA INDEX NAME)

342641-93-4 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-6-[3-(1-methylethyl)phenyl]-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]-(9CI) (CA INDEX NAME)

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342641-94-5 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl]-5-[(5-fluoro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

RN 342641-95-6 CAPLUS
CN 1H-Indole-6-carboxylic acid,
3-[4-[[2-(diethylamino)ethyl]amino]carbonyl
-3.5-dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo- (9CI) (CA
INDEX NAME)

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) pyridinylamino)sulfonyl]-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

342642-01-7 CAPLUS
1H-Pyrrole-3-carboxamide, N-[2-(diethylamino)ethyl}-5-[[5-

[(dimethylamino)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

342642-02-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[{5-[{(3-chlorophenyl)amino)sulfonyl}-1,2-dihydro-2-oxo-3H-indol-3-ylidene}methyl}-N-[2-(diethylamino)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

342641-96-7 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[5-[(dimethylamino)sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]-(9CI) (CA INDEX NAME)

342641-97-8 CAPLUS
1H-Pyrrole-3-carboxamide, 5-[[5-[[(3-chlorophenyl)amino]sulfonyl]-1,2-dhydro-2-oxo-3H-indol-3-ylidene]methyl]-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)

342641-98-9 CAPLUS 1H-Pyrrole-3-carboxamide, 5-[[1,2-dihydro-2-oxo-5-[(3-

ANSWER 63 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 342642-09-5 CAPLUS Glycine, N-[15-[1,2-dihydro-4-methyl-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-yildene|methyl]-4-methyl-1H-pyrrol-3-yilcarbonyl]-, ethyl ester (9CI) (CA INDEX NAME)

342642-10-8 CAPLUS Glycine, N-[[5-[[1,2-dihydro-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3-ylidenejmethyl]-4-methyl-H-pyrrol-3-yl|carbonyl)-, ethyl ester (9CI)

INDEX NAME)

342642-11-9 CAPLUS
Glycine, N-[[5-[{1,2-dihydro-5-[(methylamino)sulfonyl]-2-oxo-3H-indol-3ylidene]methyl]-4-methyl-1H-pyrrol-3-yl]carbonyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 64 OF 65
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INVENTOR(S):
Tang, Peng Cho; Sun, Li; Mcmahon, Gerald; Miller, Todd INVENTOR (S):

Anthony; Shirazian, Shahrzad; Wei, Chung Chen; G. Davis; Xiaoyuan, Li; Liang, Congxin Sugen, Inc., USA
PCT Int. Appl., 245 pp.
CODEN: PIXXD2
Patent
English
1

PATENT ASSIGNEE(S): SOURCE:

DOCUMENT TYPE:

DOCUMENT TIPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

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| WO 2000      | 056709      | A1 2     | 20000928 | WO 2000-US7704                                      |                 |
| W:           | AE, AG, AL, | AM, AT,  | AU, AZ,  | BA, BB, BG, BR, BY,                                 | CA, CH, CN, CR, |
|              | CU, CZ, DE, | DK, DM,  | DZ, EE,  | ES, FI, GB, GD, GE,                                 | 3H, GM, HR, HU, |
|              |             |          |          | KP, KR, KZ, LC, LK,                                 |                 |
|              | LV, MA, MD, | MG, MK,  | MN, MW,  | MX, NO, NZ, PL, PT, 1                               | RO, RU, SD, SE, |
|              | SG, SI, SK, | SL, TJ,  | TM, TR,  | TT, TZ, UA, UG, US, U                               | JZ, VN, YU, ZA, |
|              |             |          |          | RU, TJ, TM  |                 |
| RW:          | GH, GM, KE, | LS, MW,  | SD, SL,  | SZ, TZ, UG, ZW, AT, 1                               | BE, CH, CY, DE, |
|              |             |          |          | IT, LU, MC, NL, PT,                                 | SE, BF, BJ, CF, |
|              | CG, CI, CM, | GA, GN,  | GW, ML,  | MR, NE, SN, TD, TG                                  |                 |
| CA 2368      | 041         | AA 2     | 20000928 | CA 2000-2368041                                     | 20000322        |
|              |             |          |          | EP 2000-916622                                      |                 |
| R:           |             |          |          | GB, GR, IT, LI, LU, I                               | NL, SE, MC, PT, |
|              | IE, SI, LT, | LV, FI,  | RO       |   |                 |
| JP 2002      | 540096      | T2 2     | 20021126 | JP 2000-606571                                      | 20000322        |
| US 6689      | 806         | B1 2     | 20040210 | US 2000-534405                                      | 20000322        |
| PRIORITY APP | LN. INFO.:  |          |          | JP 2000-606571<br>US 2000-534405<br>US 1999-125945P | P 19990324      |
|              |             |          |          | US 1999-127863P                                     | P 19990405      |
|              |             |          |          | US 1999-131192P                                     | P 19990426      |
|              |             |          |          | US 1999-132243P                                     | P 19990503      |
|              |             |          |          | WO 2000-US7704                                      | w 20000322      |
| OTHER SOURCE | (S):        | MARPAT : | 133:2523 | 06  |                 |

ANSWER 64 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)
295799-29-0 CAPLUS
2H-Indol-2-one, 3-{[4-[3-{dimethylamino}propyl]-3,5-dimethyl-1H-pyrrol-2yl]methyl=nel-1,3-dihydro-4-{2-{3-(1-methylethyl)phenoxy}ethyl}- (9CI)
(CA INDEX NAME)

295799-47-2 CAPLUS
2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2yl]methylene]-1,3-dihydro-6-[3-pyridinyl)- (9CI) (CA INDEX NAME)

295799-51-8 CAPLUS 2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yllmethylene]-1,3-dihydro-6-(2-thienyl)- (9CI) (CA INDEX NAME)

251336-61-3P 251356-63-5P 251356-63-7P
251356-66-8P 251356-67-9P 251356-69-0P
RL: RCT (Reactant): SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (preparation of indolinones as protein kinase inhibitors)
251356-61-3 CAPLUS

L4 ANSWER 64 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

Title compds., e.g. {I; m, n = 0, 1; Q = (JR11)m; Q1 = (DR6)n; when n =

then A, B, D, E, F = C, N;  $\le 3$  of A, B, D, E, F = N; when m = 1, then G, H, J, K, L = C, N;  $\ge 1$  and  $\le 3$  of G, H, J, K, L = N; when n = 0, then A = C, N, B, F = C, N, NN, O, S; E = C, N, O, S; when m

I

O, then G = C, N, H, K, l = C, N, NH, O, S; Rl-R13 = H, alkyl, trihaloalkyl, cycloalkyl, alkenyl, alkynyl, aryl, heteroaryl, OH, alkoxy, SH, alkylthiol, aryloxy, amino, etc.; R4R5 or R5R6 or R6R7 or R7R8 =

atoms
to form a 5-6 membered (hetero)aryl ring; with addnl. provisos), were prepared Thus, 6-pyridin-3-yl-1,3-dihydroindol-2-one (preparation given),
4-methoxy-3-thien-2-ylbenzaldehyde, and piperidine were refluxed overnight
in EtOH to give 15% 3-(4-methoxy-3-thien-2-ylbenzylidene)-6-pyridin-3-yl1,3-dihydroindol-2-one. Tested title compds. inhibited HER2 kinase with ICSO = 16.4 µM to 2100 µM.

IT 251356-74-89 295799-29-0P 295799-47-2P
295799-51-89
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SFN (Synthetic preparation); THU (Therapeutic use);

logical study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (preparation of indolinones as protein kinase inhibitors) 251356-74-8 CAPLUS 2H-Indol-2-one, 3-[(4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-4-(2-hydroxyethyl)- (9CI) (CA INDEX NAME)

ANSWER 64 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 2H-Indol-2-one, 3-[[4-[3-(dimethylamino]propyl]-3,5-dimethyl-1H-pyrrol-2-yllmethylenej-1,3-dihydro-(9CI) (CA INDEX NAME)

251356-63-5 CAPLUS 2H-Indol-2-one, 5-bromo-3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-lH-pyrorl-2-yl]methylenej-1,3-dihydro- [9CI] (CA INDEX NAME)

251356-65-7 CAPLUS
2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-6-(2-methoxyphenyl)- (9CI) (CA INDEX NAME)

251356-66-8 CAPLUS
2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-6-(3-methoxyphenyl)- (9CI) (CA INDEX NAME)

251356-67-9 CAPLUS
2H-Indol-2-one, 3-{[4-{3-(dimethylamino)propyl)-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-6-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)

Page 137

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L4 ANSWER 64 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

251356-68-0 CAPLUS
2H-Indol-2-one, 5-chloro-3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1Hpyrrol-2-yl|methylene]-1,3-dihydro- (9CI) (CA INDEX NAME)

REFERENCE COUNT: THIS

31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

ANSWER 65 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

The present invention relates to 5-(2-oxo-1,2-dihydroindol-3-ylidenemethyl)-1H-pyrrol-3-ylalkanoic acid derivs. (I) (where R1 and R7 = independently H, (cyclo)alkyl, alkenyl, alkynyl, aryl, OH, alkoxy, carboxy, acetyl, (thio)amido, (trihalomethane)sultonyl, etc.; R2 = H, halo, (cyclo)alkyl, (hetero)aryl, or heteroalicyclic; R3, R4, R5, R6, R8, R9, R10 = independently H, (cyclo)alkyl, trihaloalkyl, alkenyl, alkynyl, (hetero)aryl(oxy), heteroalicyclic, OH, alkoxy, SH, alkythio, arylthio, sulfinyl, sulfonyl, sulfonamido, carbonyl, carboxy, amido, CN, NO2,

(thio)carbamyl, (un)substituted amino, etc.) which modulate the activity of protein kinases and are useful in the prevention and treatment of protein kinase related cellular disorders, such as cancer. Thus, 2,4-dimethyl-5-ethoxycarbonyl-3-(2-ethoxycarbonylethyl)pyrrole was deprotected using NaOH to form 3-(2-carboxyethyl)-2,4-dimethylpyrrole (1001) and the product C-5 formylated (two methods given for 864 and 903 yield, resp.). Reaction with 2-oxindole in EtOH and pyrrolidine or in

aqueous NaOH yielded II (88% and 91%, resp.), which reduced the average size of

NaOH yielded II (88% and 91%, resp.), which reduced the average size of
human glioma and melanoma tumors s.c. implanted in mice by 80-85%. II,
when administered orally, demonstrated notably superior efficacy compared
to structurally similar analogs.

IZ 251356-61-3P 251356-63-9P 251356-64-6P
251356-63-PP 251356-69-PP 251356-67-0P
251356-78-PP 251356-79-1P 251356-70-4P
251356-78-9P 251356-79-2P 251356-78-0P
251356-77-1P 251356-79-2P 251356-78-0P
251356-77-1P 251356-79-PP
(Biological activity or effector, except adverse); BSU
(Biological study), unclassified); SFN (Synthetic preparation); THU (Therapeutic use);
BIOL (Biological study); PREP (Preparation); USES (Uses)
(target compound; preparation of
5-(2-oxo-1,2-dihydroindol-3-ylidenemethyl)-1Hpyrrol-3-ylalkanoic acid protein kinase inhibitors as antitumor agents)

L4 ANSWER 65 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 1999:764021 CAPLUS
DOCUMENT NUMBER: 132:12257
Preparation of pyrrole substituted 2-indolinone protein kinase inhibitors
INVENTOR(S): Tang, Peng Cho: Sun, Li; McMahon, Gerald
SUNRCE: Sugen, Inc., USA
PCT Int. Appl., 240 pp.
CODEN: PIXKD2
DOCUMENT TYPE: Patent
LANGUAGE: Patent
English
English
English

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

OTHER SOURCE(S):

|       | PATENT NO. |      |       |      |     | KIND |     | DATE     |      | APPLICATION NO. |    |                                  |      |      |     |      | DATE |     |  |  |
|-------|------------|------|-------|------|-----|------|-----|----------|------|-----------------|----|----------------------------------|------|------|-----|------|------|-----|--|--|
| ,     |            |      | 61422 |      |     | A1   |     | 19991202 |      |                 |    |                                  |      |      |     |      |      |     |  |  |
|       |            | W:   | AL,   | AM,  | AT, | AU,  | AZ, | BA,      | BB,  | BG,             | BP | , BY,                            | CA,  | CH,  | CN, | CU,  | CZ,  | DE, |  |  |
|       |            |      | DK,   | EE,  | ES, | FI,  | GB, | GD,      | GE,  | GH,             | G) | 1, HR,                           | HU,  | ID,  | IL, | IN,  | IS,  | JP, |  |  |
|       |            |      |       |      |     |      |     |          |      |                 |    | , LT,                            |      |      |     |      |      |     |  |  |
|       |            |      | MW,   | MX,  | NO, | NZ,  | PL, | PT,      | RO,  | RU,             | SE | , SE,                            | SG,  | SI,  | SK, | SL,  | TJ,  | TM, |  |  |
|       |            |      |       |      |     |      |     |          |      |                 |    | (, AZ,                           |      |      |     |      |      |     |  |  |
| TM    |            |      |       |      |     |      |     |          |      |                 |    |                                  |      |      |     |      |      |     |  |  |
|       |            | RW:  | GH,   | GΜ,  | KE, | LS,  | MW, | SD,      | SL,  | SZ,             | UG | , ZW,                            | AT,  | BE,  | CH, | CY,  | DE,  | DK, |  |  |
|       |            |      |       |      |     |      |     |          |      |                 |    | , NL,                            |      |      | BF, | ВJ,  | CF,  | ÇG, |  |  |
|       |            |      | CI,   | CM,  | GΑ, | GN,  | G₩, | ML,      | MR,  | NE,             | SN | , TD,                            | TG   |      |     |      |      |     |  |  |
|       | CA         | 2314 | 156   |      |     | AA   |     | 1999     | 1202 |                 | CA | 1999-                            | 2314 | 156  |     | 1    | 9990 | 528 |  |  |
|       | ΑU         | 9944 | 102   |      |     | A1   |     | 1999     | 1213 |                 | ΑU | 1999-<br>1999-                   | 4410 | 2    |     | 1    | 9990 | 528 |  |  |
|       | ΑU         | 7592 | 26    |      |     | В2   |     | 2003     | 0410 |                 |    |                                  |      |      |     |      |      |     |  |  |
| 1     |            |      |       |      |     |      |     |          |      |                 |    | 1999-                            |      |      |     |      |      |     |  |  |
|       |            | R:   | ΑT,   | ΒE,  | CH, | DΕ,  | DK, | ES,      | FR,  | GB,             | GF | , IT,                            | LI,  | LU,  | ΝL, | SE,  | MC,  | PΤ, |  |  |
|       |            |      | ΙE,   | SI,  | LT, | LV,  | FI, | RO       |      |                 |    |                                  |      |      |     |      |      |     |  |  |
| 1     | BR         | 9910 | 792   |      |     | A_   |     | 2002     | 0129 |                 | BR | 1999-<br>2000-<br>1999-<br>2000- | 1079 | 2    |     | 1    | 9990 | 528 |  |  |
|       | TR.        | 2000 | 0351  | 4    |     | T2   |     | 2002     | 0521 |                 | TR | 2000-                            | 2000 | 0351 | 4   | 1    | 9990 | 528 |  |  |
| 1     | US         | 6395 | 734   |      |     | В1   |     | 2002     | 0528 |                 | US | 1999-                            | 3222 | 97   |     | 1    | 9990 | 528 |  |  |
|       | JP         | 2002 | 5163  | 10   |     | T2   |     | 2002     | 0604 |                 | JP | 2000-                            | 5508 | 28   |     | 1    | 9990 | 528 |  |  |
|       | NO         | 2000 | 0059  | 16   |     | A.   |     | 2001     | 129  |                 | МО | 2000-<br>2002-<br>1998-          | 5916 | _    |     | 2    | 0001 | 122 |  |  |
|       | US         | 2003 | 1051  | 51   |     | Al   |     | 2003     | 0605 |                 | US | 2002-                            | 8114 | 7_   |     | . 2  | 0020 | 225 |  |  |
| PRIOR | ITY        | APP  | LN.   | INFO | .:  |      |     |          |      |                 | us | 1998-                            | 8731 | 0P   |     | P 1  | 9980 | 529 |  |  |
|       |            |      |       |      |     |      |     |          |      |                 |    |                                  |      |      |     |      |      |     |  |  |
|       |            |      |       |      |     |      |     |          |      |                 | US | 1999-                            | 1161 | 06P  |     | P 1  | 9990 | 115 |  |  |
|       |            |      |       |      |     |      |     |          |      |                 | US | 1999-                            | 3222 | 97   |     | A1 1 | 9990 | 528 |  |  |
|       |            |      |       |      |     |      |     |          |      |                 |    |                                  |      |      |     |      |      |     |  |  |

MARPAT 132:12257

ANSWER 65 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 251356-61-3 CAPLUS 251356-61-3 CAPLUS 2H-Indol-2-one, 3-[(4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro- (9CI) (CA INDEX NAME)

251356-63-5 CAPLUS
2H-Indol-2-one, 5-bromo-3-[{4-(3-(dimethylamino)propyl)-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro- (9CI) (CA INDEX NAME)

251356-64-6 CAPLUS
2H-Indol-2-one, 3-[[4-[3-{dimethylamino}propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-6-phenyl- (9CI) (CA INDEX NAME)

251356-65-7 CAPLUS
2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-6-(2-methoxyphenyl)- [9CI) (CA INDEX NAME)

251356-66-8 CAPLUS 2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-

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ANSWER 65 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) yl]methylene]-1,3-dihydro-6-(3-methoxyphenyl)- (9CI) (CA INDEX NAME)

251356-67-9 CAPLUS
2H-Indol-2-one, 3-[{4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-6-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)

251356-68-0 CAPLUS
2H-Indol-2-one, 5-chloro-3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1Hpyrrol-2-yl]methylenej-1,3-dihydro- (9CI) (CA INDEX NAME)

251356-69-1 CAPLUS 2H-Indol-2-one, 6-chloro-3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-lH-pyrrol-2-yl]methylene]-1,3-dihydro- (9CI) (CA INDEX NAME)

ANSWER 65 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN

251356-74-8 CAPLUS
2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2yl]methylenej-1,3-dihydro-4-(2-hydroxysthyl)- (9CI) (CA INDEX NAME)

251356-75-9 CAPLUS
1H-Indole-5-sulfonamide, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1Hpyrrol-2-yl]amethylenej-2,3-dihydro-2-oxo- (9CI) (CA INDEX NAME)

251356-76-0 CAPLUS lH-Indole-5-sulfonamide,  $3-[\{4-[3-\{dimethylamino\}propyl]-3,5-dimethyl-1H-pyrcol-2-yl]methylene]-2,3-dihydro-N-(1-methylethyl)-2-oxo- (9CI) (CA INDEX NAME)$ 

251356-77-1 CAPLUS Morpholine, 4-[3-[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-2-oxo-1H-indol-5-yl]sulfonyl]- (9CI) (CA INDEX

ANSWER 65 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 251356-70-4 CAPLUS 251356-70-4 CAPLUS 2H-Indol-2-cne, 3-[(4-[3-(dimethylamino)propyl)-3,5-dimethyl-1H-pyrrol-2-yl)methylene)-1,3-dihydro-5-methoxy- (9CI) (CA INDEX NAME)

251356-71-5 CAPLUS 2H-Indol-2-one, 3-[(4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-6-methoxy- (9CI) (CA INDEX NAME)

251356-72-6 CAPLUS
2H-Indol-2-one, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-1,3-dihydro-5-methyl- (9CI) (CA INDEX NAME)

251356-73-7 CAPLUS
2H-Indol-Z-one, 3-{(4-{3-(dimethylamino)propyl)-3,5-dimethyl-1H-pyrrol-2-yllmethylene)-1,3-dihydro-4-methyl- (9CI) (CA INDEX NAME)

ANSWER 65 OF 65 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) 251356-78-2 CAPLUS | H-Indol-6-5-sulfonamide, 3-[[4-[3-(dimethylamino)propyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene]-2,3-dihydro-N,N-dimethyl-2-oxo- (9CI) (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

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LOGOFF? (Y)/N/HOLD:Y

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SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST 324.70 489.00

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION

CA SUBSCRIBER PRICE -47.45 -47.45

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